```
FILE 'HOME' ENTERED AT 10:10:51 ON 08 JAN 2003
=> fil medl capl biosis uspatf
=> s ketogenic diet
          871 KETOGENIC DIET
=> hyperuricem?
HYPERURICEM? IS NOT A RECOGNIZED COMMAND
The previous command name entered was not recognized by the system.
For a list of commands available to you in the current file, enter
"HELP COMMANDS" at an arrow prompt (=>).
=> s hyperuricem?
L2
         5411 HYPERURICEM?
=> s 11 and 12
            6 L1 AND L2
L3
=> d
     ANSWER 1 OF 6
                       MEDLINE
     2002253365
                   MEDLINE
AN
DN
     21988602 PubMed ID: 11992756
TΙ
     The ketogenic diet: a review of the experience at
     Connecticut Children's Medical Center.
     DiMario Francis J Jr; Holland Jessica
     Department of Pediatrics, University of Connecticut School of Medicine,
CS
     Division of Pediatric Neurology at Connecticut Children's Medical Center,
     Hartford 06106, USA.
so
     PEDIATRIC NEUROLOGY, (2002 Apr) 26 (4) 288-92.
     Journal code: 8508183. ISSN: 0887-8994.
CY
     United States
DТ
     Journal; Article; (JOURNAL ARTICLE)
LΑ
     English
FS
     Priority Journals
EΜ
     200207
ED
     Entered STN: 20020507
     Last Updated on STN: 20020702
     Entered Medline: 20020701
=> dup rem 13
PROCESSING COMPLETED FOR L3
              6 DUP REM L3 (0 DUPLICATES REMOVED)
=> d ibib abs 4-6
   ANSWER 4 OF 6 BIOSIS COPYRIGHT 2003 BIOLOGICAL ABSTRACTS INC.
ACCESSION NUMBER:
                    2002:619842 BIOSIS
DOCUMENT NUMBER:
                    PREV200200619842
TITLE:
                    Early- and late-onset complications of ketogenic
                    diet in intractable epilepsy.
AUTHOR (S):
                    Kang, H. C. (1); Chung, D. E. (1); Kim, H. D. (1)
CORPORATE SOURCE:
                    (1) Pediatrics and Epilepsy Center, Sang-gye Paik Hospital,
                    Seoul, Seoul South Korea
SOURCE:
                    Epilepsia, (2002) Vol. 43, No. Supplement 7, pp. 214.
                    http://blackwellscience.com/epi. print.
                    Meeting Info.: Annual Meeting of the American Epilepsy
                    Society Seattle, Washington, USA December 06-11, 2002
                    American Epilepsy Society
                    . ISSN: 0013-9580.
DOCUMENT TYPE:
                    Conference
LANGUAGE:
                    English
   ANSWER 5 OF 6 USPATFULL
ACCESSION NUMBER:
                       2001:188226 USPATFULL
TITLE:
                        DIETETIC FOOD COMPOSITION AND DIETETIC METHOD USING
                        SUCH COMPOSITION
INVENTOR(S):
                        ZOHOUNGBOGBO, MATHIAS CHRISTIAN, RIVALTA DI TORINO,
                        Italy
```

NUMBER KIND DATE

PATENT INFORMATION:

US 2001033856 A1 20011025 US 6322826 B2 20011127 US 1999-333097 A1 19990615 (9)

APPLICATION INFO.:

RELATED APPLN. INFO.: Continuation-in-part of Ser. No. US 1999-225819, filed

on 5 Jan 1999, ABANDONED

NUMBER

EP 1998-830365 19980616 PRIORITY INFORMATION:

EP 1999-201794

DOCUMENT TYPE: Utility

FILE SEGMENT: APPLICATION

LEGAL REPRESENTATIVE: SOFER & HAROUN LLP, 342 MADISON AVENUE, SUITE 1921, NEW

YORK, NY, 10173

NUMBER OF CLAIMS: 42 EXEMPLARY CLAIM: 1 LINE COUNT: 833

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

Food composition in the form of a flour comprising at least 50% of protein, less than 15% of carbohydrates and 35 to 50% of plant fibers; preferably the carbohydrate content is less than 10%, advantageously less than 5%; this composition may be used as a substitute for wheat flour in the preparation of foods such as pasta, bread, bread sticks, bakery products and pastries and constitutes the basis of a method for improving the appearance of a person by achieving a loss of weight which is beneficial from the aesthetic point of view.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

ANSWER 6 OF 6 MEDLINE

ACCESSION NUMBER: 80025910 MEDLINE

DOCUMENT NUMBER: PubMed ID: 488876 80025910

TITLE: [Possibilities for weight reduction by means of diet].

Moglichkeiten zur Verminderung des Korpergewichts mittels

diatetischer Massnahmen.

AUTHOR: Forster H

SOURCE: FORTSCHRITTE DER MEDIZIN, (1979 Aug 23) 97 (32) 1339-44.

Journal code: 2984763R. ISSN: 0015-8178. GERMANY, WEST: Germany, Federal Republic of

PUB. COUNTRY: DOCUMENT TYPE: Journal; Article; (JOURNAL ARTICLE)

LANGUAGE: German

FILE SEGMENT: Priority Journals

ENTRY MONTH: 197912

ENTRY DATE: Entered STN: 19900315

Last Updated on STN: 19900315 Entered Medline: 19791218

The different dietetic measures for weight reduction are described. According to the existing overweight the therapeutic measures are classified in four steps. In the first step, with low overweight, the energy-containing drinks (soft drinks and alcoholic beverages) are avoided. If the overweight is greater an additional reduction of the energy content of meal is required. A real reduction-diet (less than 1.000 Kcal/day or 4.200 KJ/day) demands extensive knowledge of food composition and greater efforts in meal composition. The availability of formula diets is considered as a relief. During starvation (or total fasting) as the step 4 of weight reduction diet, an extreme metabolic alteration takes place, which is characterized by ketosis. The same metabolic alteration is found by a fat-protein-diet (a so-called ketogenic diet), where hypercholesterolemia and hyperuricemia are common side effects. The carbohydrate-protein weight reduction diet is poor in health risks. Furthermore the normal metabolic pattern is maintained during this kind of diet if enough carbohydrates are provided per day (i.e. 80-100 g/day).

```
=> s hydrocoleret?
```

3 HYDROCOLERET?

=> d

ANSWER 1 OF 3 USPATFULL L5 2002:227711 USPATFULL ΑN

```
Dietetic food composition and dietetic method using such composition
ΤI
       Zohoungbogbo, Mathias C., Torino, ITALY
TN
PΙ
       US 2002122862
                        A1 20020905
       US 2001-982533
                         A1
                               20011018 (9)
ΑI
       Continuation-in-part of Ser. No. US 1999-333097, filed on 15 Jun 1999,
RLI
      GRANTED, Pat. No. US 6322826
                        19980616
PRAI
      EP 1998-830365
       EP 1999-201794
                          19990604
DT
      Utility
      APPLICATION
FS
LN.CNT 576
INCL
       INCLM: 426/549.000
NCL
      NCLM: 426/549.000
TC
      [7]
       ICM: A21D010-00
=> d 3
L5
    ANSWER 3 OF 3 USPATFULL
AN
       2001:188226 USPATFULL
ΤI
       DIETETIC FOOD COMPOSITION AND DIETETIC METHOD USING SUCH COMPOSITION
      ZOHOUNGBOGBO, MATHIAS CHRISTIAN, RIVALTA DI TORINO, Italy
IN
      US 2001033856
                        A1 20011025
                          B2
      US 6322826
                               20011127
       US 1999-333097
                         A1
                               19990615 (9)
      Continuation-in-part of Ser. No. US 1999-225819, filed on 5 Jan 1999,
RLI
      ABANDONED
PRAI
      EP 1998-830365
                          19980616
                         19990604
      EP 1999-201794
      Utility
FS
      APPLICATION
LN.CNT 833
INCL
      INCLM: 424/439.000
       INCLS: 426/557.000
      NCLM: 426/002.000
NCLS: 426/549.000; 426/601.000; 426/804.000; 514/386.000; 514/561.000
NCL
IC
      [7]
       ICM: A61K047-00
       ICS: A23P001-12
CAS INDEXING IS AVAILABLE FOR THIS PATENT.
=> fil stng
COST IN U.S. DOLLARS
                                                 SINCE FILE
                                                                 TOTAL
                                                      ENTRY
                                                              SESSION
FULL ESTIMATED COST
                                                      24.45
                                                                 25.50
FILE 'STNGUIDE' ENTERED AT 10:22:49 ON 08 JAN 2003
USE IS SUBJECT TO THE TERMS OF YOUR CUSTOMER AGREEMENT
COPYRIGHT (C) 2003 AMERICAN CHEMICAL SOCIETY, JAPAN SCIENCE
AND TECHNOLOGY CORPORATION, AND FACHINFORMATIONSZENTRUM KARLSRUHE
FILE CONTAINS CURRENT INFORMATION.
LAST RELOADED: Dec 20, 2002 (20021220/UP).
=> FIL MEDL CAPL BIOSIS USPATF
COST IN U.S. DOLLARS
                                                 SINCE FILE
                                                                 TOTAL
                                                      ENTRY
                                                              SESSION
FULL ESTIMATED COST
                                                       0.66
                                                                 26.16
FILE 'MEDLINE' ENTERED AT 10:29:16 ON 08 JAN 2003
FILE 'CAPLUS' ENTERED AT 10:29:16 ON 08 JAN 2003
USE IS SUBJECT TO THE TERMS OF YOUR STN CUSTOMER AGREEMENT.
PLEASE SEE "HELP USAGETERMS" FOR DETAILS.
COPYRIGHT (C) 2003 AMERICAN CHEMICAL SOCIETY (ACS)
FILE 'BIOSIS' ENTERED AT 10:29:16 ON 08 JAN 2003
COPYRIGHT (C) 2003 BIOLOGICAL ABSTRACTS INC. (R)
FILE 'USPATFULL' ENTERED AT 10:29:16 ON 08 JAN 2003
```

CA INDEXING COPYRIGHT (C) 2003 AMERICAN CHEMICAL SOCIETY (ACS)

```
=> s centella asiatica
L6
           655 CENTELLA ASIATICA
=> s hyperuricem? or hypouricem?
         6134 HYPERURICEM? OR HYPOURICEM?
=> s 16 and 17
             3 L6 AND L7
L8
=> dup rem 18
PROCESSING COMPLETED FOR L8
              3 DUP REM L8 (0 DUPLICATES REMOVED)
=> d tot
    ANSWER 1 OF 3 USPATFULL
AN
       2002:227711 USPATFULL
ΤI
       Dietetic food composition and dietetic method using such composition
IN
       Zohoungbogbo, Mathias C., Torino, ITALY
                             20020905
PΙ
       US 2002122862
                         A1
ΑI
       US 2001-982533
                         A1
                               20011018 (9)
       Continuation-in-part of Ser. No. US 1999-333097, filed on 15 Jun 1999,
RLI
       GRANTED, Pat. No. US 6322826
PRAI
       EP 1998-830365
                          19980616
       EP 1999-201794
                           19990604
       Utility
FS
       APPLICATION
LN.CNT 576
INCL
       INCLM: 426/549.000
NCL
       NCLM: 426/549.000
IC
       [7]
       ICM: A21D010-00
L9
    ANSWER 2 OF 3 USPATFULL
AN
       2002:185267 USPATFULL
       Dietetic food composition and dietetic method using such composition
TΤ
IN
       Zohoungbogbo, Mathias C., Torino, ITALY
                         A1 20020725
PΤ
       US 2002098175
       US 2001-982554
                               20011018 (9)
ΑI
                          A1
       Continuation-in-part of Ser. No. US 1999-333097, filed on 15 Jun 1999,
RLI
       PATENTED Continuation-in-part of Ser. No. US 1999-225819, filed on 5 Jan
       1999, ABANDONED
PRAI
       EP 1998-830365
                           19980616
       EP 1999-201794
                           19990604
DT
       Utility
FS
       APPLICATION
LN.CNT 709
       INCLM: 424/094:210
       INCLS: 514/171.000; 514/033.000; 514/540.000; 514/629.000
NCL
       NCLM: 424/094.210
       NCLS: 514/171.000; 514/033.000; 514/540.000; 514/629.000
IC
       [7]
       ICM: A61K038-54
       ICS: A61K031-704; A61K031-56; A61K031-16; A61K031-24
CAS INDEXING IS AVAILABLE FOR THIS PATENT.
    ANSWER 3 OF 3 USPATFULL
L9
       2001:188226 USPATFULL
AN
TI
       DIETETIC FOOD COMPOSITION AND DIETETIC METHOD USING SUCH COMPOSITION
       ZOHOUNGBOGBO, MATHIAS CHRISTIAN, RIVALTA DI TORINO, Italy
IN
PΙ
       US 2001033856
                          A1
                               20011025
       US 6322826
                          B2
                               20011127
       US 1999-333097
AΙ
                         A1
                               19990615 (9)
RLI
       Continuation-in-part of Ser. No. US 1999-225819, filed on 5 Jan 1999,
       ABANDONED
PRAI
       EP 1998-830365
                           19980616
       EP 1999-201794
                           19990604
DT
       Utility
      APPLICATION
LN.CNT 833
INCL
       INCLM: 424/439.000
       INCLS: 426/557.000
```

NCLM: 426/002.000 NCL

NCLS: 426/549.000; 426/601.000; 426/804.000; 514/386.000; 514/561.000

IC [7]

> ICM: A61K047-00 ICS: A23P001-12

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

=> s benfluorex

L10 276 BENFLUOREX

=> s hypercholesterolemia

47365 HYPERCHOLESTEROLEMIA

=> s 110 and 111

33 L10 AND L11 L12

=> dup rem 112

PROCESSING COMPLETED FOR L12

29 DUP REM L12 (4 DUPLICATES REMOVED)

=> d ibib abs 26-29

L13 ANSWER 26 OF 29 MEDLINE DUPLICATE 2

ACCESSION NUMBER: 1998330340 MEDLINE

DOCUMENT NUMBER: 98330340 PubMed ID: 9667766

TITLE:

Metabolic and anti-atherogenic effects of long-term benfluorex in dyslipidemic insulin-resistant sand

rats (Psammomys obesus).

Marquie G; El Madani T; Solera M L; Pieraggi M T; Hadjiisky AUTHOR:

P; Ravel D; Seguin L; Bennani N

Laboratoire de Recherche des Macrophages, Mediateurs de CORPORATE SOURCE:

l'Inflammation et Interactions Cellulaires, Toulouse,

France.

SOURCE: LIFE SCIENCES, (1998) 63 (1) 65-76. Journal code: 0375521. ISSN: 0024-3205.

ENGLAND: United Kingdom

PUB. COUNTRY: DOCUMENT TYPE: Journal; Article; (JOURNAL ARTICLE)

LANGUAGE: English

FILE SEGMENT: Priority Journals

ENTRY MONTH: 199807

ENTRY DATE: Entered STN: 19980731

Last Updated on STN: 19980731

Entered Medline: 19980723

Benfluorex is a clinical lipid-lowering agent with

antihyperglycemic properties. The effect of long-term oral treatment (10 mg/kg/day for 7.5 months) on carbohydrate and lipid metabolism and aortic morphology was investigated in 24 insulin-resistant sand rats receiving a standard laboratory diet supplemented with cholesterol (2%). Untreated controls (n=34) developed impaired glucose tolerance, hyperinsulinemia, hypertriglyceridemia, hypercholesterolemia and elevated plasma LDL- and VLDL-cholesterol, positively correlated with the proportion of the thoracic aorta displaying oil red O-positive atherosclerosis; ultrastructural examination showed intimal lipid deposits, foam cells, polymorph infiltrates and fibrosis. Benfluorex-treated animals showed significant decreases in glucose intolerance, hyperinsulinemia, hypertriglyceridemia, hypercholesterolemia, and plasma LDL- and VLDL-cholesterol, with no evidence of aortic atheroma. The metabolic benefits of benfluorex may protect against the long-term development of atherosclerosis in the insulin-resistant dyslipidemic syndrome.

L13 ANSWER 27 OF 29 MEDIATNE

ACCESSION NUMBER: 93066063 MEDITNE

DOCUMENT NUMBER: 93066063 PubMed ID: 1438102

TITLE: [Mode of action of benfluorex. Recent data].

Mode d'action du benfluorex. Donnees recentes.

AUTHOR: Brindley D N

CORPORATE SOURCE: Department of Biochemistry, University of Alberta,

Edmonton, Canada.

SOURCE: PRESSE MEDICALE, (1992 Sep 9) 21 (28) 1330-5. Ref: 26

Journal code: 8302490. ISSN: 0755-4982.

PUB. COUNTRY: France DOCUMENT TYPE: Journal; Article; (JOURNAL ARTICLE)

General Review; (REVIEW)

(REVIEW, TUTORIAL)

LANGUAGE: French

FILE SEGMENT: Priority Journals

ENTRY MONTH:

ENTRY DATE: Entered STN: 19930122

199212

Last Updated on STN: 19930122 Entered Medline: 19921214

An increased risk of developing premature atherosclerosis is associated with stress, diabetes, obesity, and hypertension. These conditions are associated with insulin resistance, hyperglycemia, hypertriglyceridemia and hypercholesterolemia. An alternative way of interpreting insulin resistance is to consider that metabolism in this condition would be regulated to a greater extent by stress hormones and in particular by cortisol. Glucocorticoids and fatty acids (which are produced in response to stress) antagonise the actions of insulin in promoting glucose uptake and protein synthesis, in decreasing gluconeogenesis and protein catabolism, and promoting the clearance of intermediate density lipoprotein and low density lipoprotein from the circulation by the liver. They also promote the secretion of very low density lipoprotein thus producing hypertriglyceridemia and hypercholesterolemia. By contrast to this antagonism, cortisol can also facilitate the action of insulin in stimulating the storage of energy via glycogen and fatty acid synthesis and through lipoprotein lipase in adipose tissue. These effects are significant in relation to obesity and to weight gain. An increased control of metabolism by cortisol therefore produces changes in metabolism that are potentially atherogenic and it is associated with insulin resistance and the other risk factors for atherosclerosis. Benfluorex treatment improves insulin sensitivity and has antihyperglycemic and hypolipidemic effects in human beings and in experimental animals. These effects can be observed independently of weight loss, but lowering food intake also produces a metabolic benefit. Long-term treatment with benfluorex can also decrease stress responses in terms of glucocorticoid release and the stimulation of lipolysis probably by its serotoninergic control of the hypothalamic-pituitary-adrenal axis. Such an action provides for an integrated treatment of the obese-diabetic-hyperlipidemic syndrome. Benfluorex produces overall changes in metabolism that tend to normalise the major risk factors associated with premature atherosclerosis. This provides a potential advantage over other therapies for atherosclerosis which may ameliorate a symptom (e.g., hyperlipidemia) without treating the underlying metabolic disturbance that predisposes to

L13 ANSWER 28 OF 29 CAPLUS COPYRIGHT 2003 ACS DUPLICATE 3

ACCESSION NUMBER: 1991:550130 CAPLUS

DOCUMENT NUMBER: 115:150130

atherogenesis.

TITLE: Decreased serum lipids, serum insulin and

triacylglycerol synthesis in adipose tissue of JCR:LA-corpulent rats treated with benfluorex

AUTHOR(S): Brindley, David N.; Hales, Paul; Al-Sieni, Abdulbasit

I. I.; Russell, James C.

CORPORATE SOURCE: Dep. Biochem., Univ. Alberta, Edmonton, AB, T6G 2S2,

Can.

SOURCE: Biochimica et Biophysica Acta (1991), 1085(1), 119-25

CODEN: BBACAQ; ISSN: 0006-3002

DOCUMENT TYPE: Journal LANGUAGE: English

AB Rats of the JCR:LA-corpulent strain were treated with benfluorex daily at a dose of 25 mg/kg. This strain of rat, if homozygous for the cp gene (cp/cp), is hyperphagous, obese, hypertriglyceridemic, insulin-resistant and in the case of male rats, atherosclerosis prone. The benfluorex treatment produced a sharp redn. in food intake which remained suppressed despite recovery toward normal after 2 wk of treatment. This was accompanied by sustained decreases in body wt. and adipose tissue mass. The ability of adipose tissue from female rats to take up glucose and convert it to lactate, glyceride-glycerol and fatty acids was decreased. This decrease was largely due to decreased adipose tissue mass. The serum concns. of glucose, lactate, triacylglycerol, cholesterol, phospholipids and insulin were decreased in both sexes. The treatment also improved glucose tolerance and decreased corticosterone concns. in male rats only. While redn. of food consumption contributes to

the effects seen, benfluorex clearly had significant direct metabolic effects. The effects are consistent with an improved insulin sensitivity leading to a decrease in circulating triacylglycerol. The changes produced by benfluorex are all in directions that should inhibit atherogenesis in this animal model for the human obesity (hypertriglyceridemia) insulin-resistant syndrome.

L13 ANSWER 29 OF 29 MEDLINE

ACCESSION NUMBER: 81103256 DOCUMENT NUMBER: 81103256

31103256 MEDLINE

DOCUMBIA

81103256 PubMed ID: 6256875

TITLE:

[Cross-over study of benfluorex and a hypolipemic agent in diet-resistant types IV and II b hyperlipemia

(author's transl)].

Etude de l'efficacite du benfluorex et d'un

hypolipidemiant de reference en traitement croise dans les hyperlipidemies de types IV et II b non corrigees par le

regime seul.

AUTHOR:

Graisely B; Cloarec M

SOURCE:

SEMAINE DES HOPITAUX, (1980 Jun 18-25) 56 (25-28) 1221-5.

Journal code: 9410059.

PUB. COUNTRY:

France

DOCUMENT TYPE:

Journal; Article; (JOURNAL ARTICLE)

LANGUAGE: French

FILE SEGMENT:

Priority Journals

ENTRY MONTH:

198103

ENTRY DATE:

Entered STN: 19900316

Last Updated on STN: 19900316

Entered Medline: 19810324

AB The effects of benfluorex and a hypolipemic agent were compared in 24 patients with types IV or II b hyperlipoproteinemia unimproved after 4 months diet, by means of a cross-over study with two periods of two months treatment separated by two months without treatment. Results showed similar efficacy for each treatment against lipid parameters; statistically significant variations in body weight (-0,8%; p < 0,0001), fasting blood sugar (-11,7%; p < 0.001), and of uric acid (-8,3%; p greater than or equal to 0,001) with benfluorex VS., respectively (-0,1%; P: NS), (+ 3,4%; P < 0.05), and (+ 1,3% PP: NS) with the hypolipemic agent; statistically significant reductions in VLDL electrophoretic levels; (29,7%: P < 0,001) with benfluorex VS., (17,3%; P < 0.001) with the hypolipemic agent; and an increase in HDL electrophoretic levels; benfluorex (10,7%; P < 0,001) VS., (0,9%; P: NS) with the hypolipemic agent.

=> FIL STNGUIDE

COST IN U.S. DOLLARS SINCE FILE TOTAL ENTRY SESSION FULL ESTIMATED COST 21.83 47.99 DISCOUNT AMOUNTS (FOR QUALIFYING ACCOUNTS) SINCE FILE TOTAL ENTRY SESSION CA SUBSCRIBER PRICE -0.65 -0.65

FILE 'STNGUIDE' ENTERED AT 10:33:38 ON 08 JAN 2003 USE IS SUBJECT TO THE TERMS OF YOUR CUSTOMER AGREEMENT COPYRIGHT (C) 2003 AMERICAN CHEMICAL SOCIETY, JAPAN SCIENCE AND TECHNOLOGY CORPORATION, AND FACHINFORMATIONSZENTRUM KARLSRUHE

FILE CONTAINS CURRENT INFORMATION. LAST RELOADED: Dec 20, 2002 (20021220/UP).

=> s metformin

0 METFORMIN

L14

0 METFORMIN

=> FIL MEDL CAPL BIOSIS USPATF

COST IN U.S. DOLLARS SINCE FILE TOTAL ENTRY SESSION FULL ESTIMATED COST 0.54 48.53 DISCOUNT AMOUNTS (FOR QUALIFYING ACCOUNTS) SINCE FILE TOTAL ENTRY SESSION CA SUBSCRIBER PRICE 0.00 -0.65

```
FILE 'MEDLINE' ENTERED AT 10:38:58 ON 08 JAN 2003
FILE 'CAPLUS' ENTERED AT 10:38:58 ON 08 JAN 2003
USE IS SUBJECT TO THE TERMS OF YOUR STN CUSTOMER AGREEMENT.
PLEASE SEE "HELP USAGETERMS" FOR DETAILS.
COPYRIGHT (C) 2003 AMERICAN CHEMICAL SOCIETY (ACS)
FILE 'BIOSIS' ENTERED AT 10:38:58 ON 08 JAN 2003
COPYRIGHT (C) 2003 BIOLOGICAL ABSTRACTS INC.(R)
FILE 'USPATFULL' ENTERED AT 10:38:58 ON 08 JAN 2003
CA INDEXING COPYRIGHT (C) 2003 AMERICAN CHEMICAL SOCIETY (ACS)
=> s metformin
L15
         5863 METFORMIN
=> d his
     (FILE 'HOME' ENTERED AT 10:10:51 ON 08 JAN 2003)
     FILE 'MEDLINE, CAPLUS, BIOSIS, USPATFULL' ENTERED AT 10:13:40 ON 08 JAN
     2003
L1
            871 S KETOGENIC DIET
L2
           5411 S HYPERURICEM?
              6 S L1 AND L2
L3
L4
              6 DUP REM L3 (0 DUPLICATES REMOVED)
              3 S HYDROCOLERET?
1.5
     FILE 'STNGUIDE' ENTERED AT 10:22:49 ON 08 JAN 2003
     FILE 'MEDLINE, CAPLUS, BIOSIS, USPATFULL' ENTERED AT 10:29:16 ON 08 JAN
     2003
L6
            655 S CENTELLA ASIATICA
           6134 S HYPERURICEM? OR HYPOURICEM?
1.7
              3 S L6 AND L7
              3 DUP REM L8 (0 DUPLICATES REMOVED)
T.9
            276 S BENFLUOREX
L10
L11
          47365 S HYPERCHOLESTEROLEMIA
L12
             33 S L10 AND L11
L13
             29 DUP REM L12 (4 DUPLICATES REMOVED)
     FILE 'STNGUIDE' ENTERED AT 10:33:38 ON 08 JAN 2003
1.14
              0 S METFORMIN
     FILE 'MEDLINE, CAPLUS, BIOSIS, USPATFULL' ENTERED AT 10:38:58 ON 08 JAN
           5863 S METFORMIN
L15
=> s 111 and 115
L16
          147 L11 AND L15
=> dup rem 116
PROCESSING COMPLETED FOR L16
           134 DUP REM L16 (13 DUPLICATES REMOVED)
L17
=> s 111 (S) 115
            23 L11 (S) L15
L18
=> dup rem 118
PROCESSING COMPLETED FOR L18
            19 DUP REM L18 (4 DUPLICATES REMOVED)
L19
=> d ibib abs 15-19
L19 ANSWER 15 OF 19
                         MEDLINE
                                                        DUPLICATE 2
ACCESSION NUMBER: 88240640
                                MEDITNE
DOCUMENT NUMBER:
                    88240640
                               PubMed ID: 3377878
TITLE:
                    The antidiabetic drug metformin decreases cholesterol
                    metabolism in cultured human fibroblasts.
AUTHOR:
                    Maziere J C; Maziere C; Mora L; Gardette J; Salmon S;
```

Auclair M; Polonovski J

Faculte de Medecine Saint-Antoine, UA 524 du CNRS, Paris,

CORPORATE SOURCE:

France.

SOURCE: ATHEROSCLEROSIS, (1988 May) 71 (1) 27-33.

Journal code: 0242543. ISSN: 0021-9150.

PUB. COUNTRY: Netherlands

DOCUMENT TYPE: Journal; Article; (JOURNAL ARTICLE)

LANGUAGE: English

FILE SEGMENT: Priority Journals ENTRY MONTH: 198806

ENTRY DATE: Entered STN: 19900308

Last Updated on STN: 19980206 Entered Medline: 19880630

The effect of the hypoglycemic biguanide drug Metformin was investigated after a 72 h pretreatment of human cultured fibroblasts. Metformin induced a moderate increase in low density lipoprotein binding, uptake and internalization (25% increase after treatment with 5 X 10(-4) M of drug). A decrease in sterol, fatty acid and triacyglycerol synthesis from sodium acetate was observed after pretreatment with the drug, with a dose-dependent effect in the range of 5 X 10(-5) to 5 X 10(-4) M (50% reduction of sterol synthesis after treatment with Metformin 5 X 10(-4) M). This effect was also observed in fibroblasts from a patient with homozygous familial hypercholesterolemia. Cholesterol esterification studied by incorporation of radiolabeled oleic acid was reduced by Metformin (40% of control after treatment with Metformin 5 X 10(-4) M) whereas incorporation into triacylglycerols was less impaired. These effects of Metformin on cholesterol metabolism were observed either in the presence or in the absence of low density lipoproteins. Moreover, Metformin also reduced cholesterol esterification in J774 monocyte-macrophage cells. Metformin also induced a decrease of hydroxymethylglutaryl coenzyme A reductase activity in cultured fibroblasts and a reduction of acyl-coenzyme A: cholesterol-O-acyltransferase activity in cultured fibroblasts and J774

L19 ANSWER 16 OF 19 CAPLUS COPYRIGHT 2003 ACS

ACCESSION NUMBER: 1983:3289 CAPLUS

DOCUMENT NUMBER: 98:3289

cells.

TITLE: Arachidonic acid metabolites in the interaction

between platelets and arterial walls

AUTHOR(S): Paoletti, R.; Tremoli, E.

CORPORATE SOURCE: Inst. Pharmacol. Pharmacogn., Univ. Milan, Milan,

Italy

SOURCE: Giornale della Arteriosclerosi (1982), 7(1), 213-16

CODEN: GIARA5; ISSN: 0017-0224

DOCUMENT TYPE: Journal LANGUAGE: English

AB Blood platelets from patients with hypercholesterolemia (HC) type IIA, stimulated in vitro with 25 units thrombin/mL or 30 .mu.g collagen/mL, showed malondialdehyde levels that were above those of similarly treated normal platelets. Blood platelets from rabbits with dietary HC were aggregated by lower concns. of collagen than normal platelets. Platelets from HC rabbits treated with metformin (a drug which does not affect blood plasma cholesterol but increases platelet membrane fluidity) showed nearly normal collagen-induced aggregability. Thus, HC in rabbits induces hyperaggregability of blood platelets which is similar to that of platelets of patients with HC type IIA.

L19 ANSWER 17 OF 19 CAPLUS COPYRIGHT 2003 ACS ACCESSION NUMBER: 1982:433261 CAPLUS

DOCUMENT NUMBER: 97:33261

TITLE: Metformin reduces platelet hypersensitivity in

hypercholesterolemic rabbits

AUTHOR(S): Tremoli, Elena; Ghiselli, Giancarlo; Maderna, Paola;

Colli, Susanna; Sirtori, Cesare R.

CORPORATE SOURCE: Inst. Pharmacol. Pharmacogn., Univ. Milan, Milan,

20129, Italy

SOURCE: Atherosclerosis (Shannon, Ireland) (1982), 41(1),

53-60

CODEN: ATHSBL; ISSN: 0021-9150

DOCUMENT TYPE: Journal LANGUAGE: English

AB The effects of metformin on platelet responsiveness to aggregating agents were studied in cholesterol-fed rabbits. Three groups of animals were

fed, for 1 mo, either a normal (N), or a hypercholesterolemic (HC), or a hypercholesterolemic plus 0.5% metformin diet (HC + Met). Platelets from the HC rabbits required significantly lower collagen and arachidonic acid concns. to aggregate, as compared to platelets from N rabbits. The platelet response from the HC + Met rabbits was not significantly different from that of normals. The cholesterol/phospholipid ratio in platelets was increased in both dietary groups (HC, HC + Met). The serum thromboxane B2 concns. did not show any significant difference between the groups. Plasma exchange expts. failed to indicate a specific effect of the plasma environment on platelet behavior. In view of the inactivity of metformin on the platelet cyclooxygenase pathway, the reported results suggest that metformin may act by an as yet unexplored mechanism.

L19 ANSWER 18 OF 19 MEDLINE DUPLICATE 3

ACCESSION NUMBER: 82027578 MEDLINE

DOCUMENT NUMBER: 82027578 PubMed ID: 7286137

TITLE: Changes in the lipoproteins of rabbits on a high-fat,

cholesterol-free diet; preventive action of metformin.

AUTHOR: Lacombe C; Nibbelink M

SOURCE: EXPERIENTIA, (1981) 37 (8) 854-5.

Journal code: 0376547. ISSN: 0014-4754.

PUB. COUNTRY: Switzerland

DOCUMENT TYPE: Journal; Article; (JOURNAL ARTICLE)

LANGUAGE: English

FILE SEGMENT: Priority Journals

ENTRY MONTH: 198112

ENTRY DATE: Entered STN: 19900316

Last Updated on STN: 19900316

Entered Medline: 19811221

AB Endogenous hypercholesterolemia induced by a cholesterol-free, high-fat diet corresponds to an increase in the level of low density lipoproteins and their enrichment in cholesterol esters. Metformin has no effect on the rise in plasma cholesterol but completely prevents the appearance of cholesterol-rich low-density lipoprotein.

L19 ANSWER 19 OF 19 CAPLUS COPYRIGHT 2003 ACS ACCESSION NUMBER: 1979:197644 CAPLUS

ACCESSION NUMBER: 1979:197644 C

DOCUMENT NUMBER: 90:197644

TITLE: Turnover and aortic uptake of very low density

lipoproteins (VLDL) from hypercholesteremic rabbits:

effect of metformin

AUTHOR(S): Sirtori, Cesare R.; Innocenti, L.; Grigolato, P. G.;

Rodriguez, J.

CORPORATE SOURCE: Inst. Morbid Anat., Univ. Milan, Milan, Italy

SOURCE: Advances in Experimental Medicine and Biology (1977),

82(Atheroscler.: Metab., Morphol., Clin. Aspects),

268-71

CODEN: AEMBAP; ISSN: 0065-2598

DOCUMENT TYPE: Journal LANGUAGE: English

AB Metformin [657-24-9] administration (135 mg/kg/day in diet, for 4 wk) only slightly modified cholesterol [57-88-5] induced hyperlipidemia (from 1403 to 978 mg/dL) in rabbits. Triglyceride and blood sugar levels were not different. Aortic and liver lipids markedly decreased and the compn. of very low d. lipoproteins (VLDL) changed in treated rabbits. Protein concn., phosphatidylethanolamines, and phosphatidylinositols increased whereas cholesterol esters and sphingomyelin content decreased. Turnover of VLDL was markedly accelerated and the uptake of the labeled VLDL into the aorta was lower in the treated animals. VLDL and low d. lipoproteins taken from metformin treated animals had lower affinity for the aortic lipoprotein complexing factor than those from control animals. Thus, metformin can markedly reduce atherosclerosis while only moderately influencing hyperlipidemia.

=> d ibib abs 11-14

AUTHOR:

L19 ANSWER 11 OF 19 MEDLINE

ACCESSION NUMBER: 2001208494 MEDLINE

DOCUMENT NUMBER: 21176757 PubMed ID: 11281188

TITLE: Myositis, microvesicular hepatitis, and progression to

cirrhosis from troglitazone added to simvastatin. Caldwell S H; Hespenheide E E; von Borstel R W

CORPORATE SOURCE: Department of Internal Medicine, University of Virginia,

Charlottesville, USA.

SOURCE: DIGESTIVE DISEASES AND SCIENCES, (2001 Feb) 46 (2) 376-8.

Journal code: 7902782. ISSN: 0163-2116.

PUB. COUNTRY: United States

DOCUMENT TYPE: Journal; Article; (JOURNAL ARTICLE)

LANGUAGE: English

FILE SEGMENT: Abridged Index Medicus Journals; Priority Journals

ENTRY MONTH: 200104

ENTRY DATE: Entered STN: 20010417

Last Updated on STN: 20010417 Entered Medline: 20010412

AB A 68-year-old woman, with type 2 diabetes mellitus,

hypercholesterolemia, and prior long-term simvastatin therapy, self-resumed troglitazone after running out of metformin. She developed an acute severe hepatitis with microvesicular steatosis and mysositis. There was subsequent resolution of the myositis but progression of the hepatitis to symptomatic cirrhosis over a period of 12 weeks. Both troglitazone and simvastatin are metabolized by cytochrome P-450 3A4. Troglitazone typically induces metabolism of drugs metabolized by this cytochrome so that simple simvastatin toxicity seems less likely to have been involved. The association with myositis, the severity of the hepatitis with progression to cirrhosis, and the presence of microvesicular steatosis suggests altered mitochondrial metabolism, which has been described with each agent, as the underlying pathogenic mechanism. Although troglitazone (Rezulin) has been withdrawn from the market, other similar agents are available for therapy of type 2 diabetes mellitus. Increased awareness of a potential interaction between these two classes of drugs is warranted.

L19 ANSWER 12 OF 19 MEDLINE

ACCESSION NUMBER: 2001303127 MEDLINE

DOCUMENT NUMBER: 20540820 PubMed ID: 11092283

TITLE: The Diabetes Prevention Program: baseline characteristics

of the randomized cohort. The Diabetes Prevention Program

Research Group.

AUTHOR: Anonymous SOURCE: DIABETES (

DIABETES CARE, (2000 Nov) 23 (11) 1619-29.

Journal code: 7805975. ISSN: 0149-5992.

PUB. COUNTRY: United States
DOCUMENT TYPE: (CLINICAL TRIAL)

Journal; Article; (JOURNAL ARTICLE)

(MULTICENTER STUDY)

(RANDOMIZED CONTROLLED TRIAL)

LANGUAGE: English

FILE SEGMENT: Priority Journals

ENTRY MONTH: 200105

ENTRY DATE: Entered STN: 20010604

Last Updated on STN: 20010604 Entered Medline: 20010531

AB OBJECTIVE: The Diabetes Prevention Program (DPP) is a 27-center randomized clinical trial designed to evaluate the safety and efficacy of interventions that may delay or prevent development of diabetes in people at increased risk for type 2 diabetes. RESEARCH DESIGN AND METHODS: Eligibility requirements were age > or = 25 years, BMI > or = 24 kg/m2 (> or = 22 kg/m2 for Asian-Americans), and impaired glucose tolerance plus a fasting plasma glucose of 5.3-6.9 mmol/l (or < or = 6.9 mmol for American Indians). Randomization of participants into the DPP over 2.7 years ended in June 1999. Baseline data for the three treatment groups--intensive lifestyle modification, standard care plus metformin, and standard care plus placebo--are presented for the 3,234 participants who have been randomized. RESULTS: Of all participants, 55% were Caucasian, 20% were African-American, 16% were Hispanic, 5% were American Indian, and 4% were Asian-American. Their average age at entry was 51 +/- 10.7 years (mean +/- SD), and 67.7% were women. Moreover, 16% were < 40 years of age, and 20% were > or = 60 years of age. Of the women, 48% were postmenopausal. Men and women had similar frequencies of history of hypercholesterolemia (37 and 33%, respectively) or hypertension (29 and 26%, respectively). On the basis of fasting lipid determinations, 54% of men and 40% of women fit National Cholesterol Education Program criteria for abnormal lipid profiles. More men than women were current or former cigarette smokers or had a history of coronary heart disease. Furthermore, 66% of men and 71% of women had a first-degree relative with

diabetes. Overall, BMI averaged 34.0 +/- 6.7 kg/m2 at baseline with 57% of the men and 73% of women having a BMI > or = 30 kg/m2. Average fasting plasma glucose (6.0 +/- 0.5 mmol/l) and HbA1c (5.9 +/- 0.5%) in men were comparable with values in women (5.9 +/- 0.4 mmol/l and 5.9 +/- 0.5%, respectively). CONCLUSIONS: The DPP has successfully randomized a large cohort of participants with a wide distribution of age, obesity, and ethnic and racial backgrounds who are at high risk for developing type 2 diabetes. The study will examine the effects of interventions on the development of diabetes.

L19 ANSWER 13 OF 19 MEDLINE

ACCESSION NUMBER: 96084542 MEDLINE

DOCUMENT NUMBER: 96084542 PubMed ID: 8557288

Special features of coronary heart disease in people of the TITLE:

Indian sub-continent.

AUTHOR: Vardan S; Mookherjee S; Vardan S; Sinha A K CORPORATE SOURCE: V.A. Medical Center, Syracuse, New York, USA.

SOURCE: INDIAN HEART JOURNAL, (1995 Jul-Aug) 47 (4) 399-407. Ref:

126 Journal code: 0374675. ISSN: 0019-4832.

PUB. COUNTRY: India

DOCUMENT TYPE: Journal; Article; (JOURNAL ARTICLE)

General Review; (REVIEW)

(REVIEW, ACADEMIC)

LANGUAGE: English

FILE SEGMENT: Priority Journals

ENTRY MONTH: 199602

Entered STN: 19960312 ENTRY DATE:

Last Updated on STN: 19960312 Entered Medline: 19960228

Contrary to the popular belief, coronary heart disease (CHD) is indeed common in the Indian sub-continent. Expatriate Indians in their newly adopted countries have 3 to 5 times more chance of developing CHD than the native population or the other immigrant groups. The well-known risk factors such as hypercholesterolemia, hypertension and smoking do not appear to play a major role, while the syndrome of insulin resistance seems to be an important risk factor for CHD in people of this sub-continent. Abdominal obesity, hypertriglyceridemia, and low plasma HDL cholesterol are the markers of this syndrome. Increased plasma insulin levels or even better, the C-peptide measurement may help in identifying the abnormality early. As CHD among Indians has been found to be severe and more diffuse with serious complications and increased mortality at a younger age, preventive measures need to be instituted early. Low fat and complex carbohydrate diet along with regular aerobic exercise may help reduce abdominal obesity, improve insulin sensitivity and HDL cholesterol levels. Hypertriglyceridemia uncontrolled by above measures may require pharmacotherapy with agents such as gemfibrozil. Smoking must be stopped to help reduce insulin resistance and improve HDL levels and endothelial function. Those with hypertension should be considered for therapy with ACE inhibitors, which may improve insulin sensitivity. In patients with insulin resistance, therapy with metformin or troglitazone may be helpful.

L19 ANSWER 14 OF 19 MEDLINE DUPLICATE 1

ACCESSION NUMBER: 91151686 MEDLINE

DOCUMENT NUMBER: 91151686 PubMed ID: 2291838

TITLE: Cholesterol lowering effect of metformin in combined

hyperlipidemia: placebo controlled double blind trial.

AUTHOR: Pentikainen P J; Voutilainen E; Aro A; Uusitupa M; Penttila

I; Vapaatalo H

Third Department of Medicine, Helsinki University Central CORPORATE SOURCE:

Hospital, Finland.

SOURCE: ANNALS OF MEDICINE, (1990) 22 (5) 307-12.

Journal code: 8906388. ISSN: 0785-3890.

PUB. COUNTRY: Finland

DOCUMENT TYPE: (CLINICAL TRIAL)

Journal; Article; (JOURNAL ARTICLE)

(RANDOMIZED CONTROLLED TRIAL)

LANGUAGE: English

FILE SEGMENT: Priority Journals

ENTRY MONTH: 199104

ENTRY DATE: Entered STN: 19910428

Last Updated on STN: 19970203

Entered Medline: 19910410

AR Metformin, an antidiabetic biguanide derivative, prevents experimental atherosclerosis and induces structural changes in lipoproteins in experimental animals. In the present study we investigated the effect of metformin on serum lipoproteins and platelet function in 24 non-diabetic patients with type II B hyperlipidemia. The patients were randomly given metformin in two dosage levels (1.0 g/day and 2.0 g/day) and placebo for periods of nine weeks in a crossover trial. Metformin caused a dose dependent fall in the concentrations of total serum cholesterol and of LDL-cholesterol. The average concentration of total cholesterol was 8.54 +/- 0.22 (SE) mmol/l, 8.12 + - 0.19 mmol/l and 7.79 + - 0.15 mmol/l during placebo, metformin 1.0 g/day and 2.0 g/day treatments, respectively. Both metformin values differed significantly (P less than 0.05) from the placebo value. Thus there was an average fall of 8.1% in total cholesterol after the higher metformin dose. LDL-cholesterol was 5.25 +/- 0.23 mmol/l after placebo, falling by 3.1% and 9.6% after metformin doses of 1.0 g/day and 2.0 g/day, respectively. The concentrations of HDL-cholesterol and total serum triglycerides showed no significant changes. Body weight, blood glucose, plasma insulin, blood lactate, platelet function and urinary excretion of prostanoids remained unchanged during the study. The reduction of total- and LDL-cholesterol levels may be a welcome additional consequence of metformin during treatment of diabetic patients with hypercholesterolemia.

=> s 16 and 111

L20 4 L6 AND L11

=> dup rem 120

PROCESSING COMPLETED FOR L20

4 DUP REM L20 (0 DUPLICATES REMOVED) L21

=> d ti tot

L21 ANSWER 1 OF 4 USPATFULL

тT Dietetic food composition and dietetic method using such composition

L21 ANSWER 2 OF 4 USPATFULL

Dietetic food composition and dietetic method using such composition

L21 ANSWER 3 OF 4 USPATFULL

TI Triterpene compositions and methods for use thereof

L21 ANSWER 4 OF 4 USPATFULL

TI DIETETIC FOOD COMPOSITION AND DIETETIC METHOD USING SUCH COMPOSITION

=> d 3 ibib abs

L21 ANSWER 3 OF 4 USPATFULL

ACCESSION NUMBER: 2002:224280 USPATFULL

TITLE: Triterpene compositions and methods for use thereof

Arntzen, Charles J., Ithaca, NY, United States INVENTOR(S): Blake, Mary E., Tucson, AZ, United States

Gutterman, Jordan U., Houston, TX, United States Hoffmann, Joseph J., Tucson, AZ, United States Jayatilake, Gamini S., Broomfield, CO, United States

Bailey, David T., Boulder, CO, United States Research Development Foundation, Carson City, NV,

PATENT ASSIGNEE(S): United States (U.S. corporation)

NUMBER KIND DATE PATENT INFORMATION: US 6444233 B1 20020903 APPLICATION INFO.: US 1999-314691 19990519 (9)

NUMBER DATE

US 1998-99066P 19980903 (60) US 1998-85997P 19980519 (60) PRIORITY INFORMATION:

DOCUMENT TYPE: Utility FILE SEGMENT: GRANTED

PRIMARY EXAMINER: Ta ASSISTANT EXAMINER: F1

Tate, Christopher R. Flood, Michele C.

LEGAL REPRESENTATIVE:

Fulbright & Jaworski LLP

NUMBER OF CLAIMS: EXEMPLARY CLAIM: 18 1

NUMBER OF DRAWINGS:

73 Drawing Figure(s); 43 Drawing Page(s)

LINE COUNT:

7526

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

AB The invention provides novel saponin mixtures and compounds which are isolated from the species Acacia victoriae and methods for their use. These compounds may contain a triterpene moiety, such as acacic or oleanolic acid, to which oligosaccharides and monoterpenoid moieties are attached. The mixtures and compounds have properties related to the regulation of apoptosis and cytotoxicity of cells and exhibit potent anti-tumor effects against a variety of tumor cells.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

=> d 3 kwic

L21 ANSWER 3 OF 4 USPATFULL

DETD The two main saponins asiaticoside and madecassoside from Centella asiatica (Umbelliferae) have been separated

with the aid of an Ito multi-layer coil separator-extractor (P.C. Inc.)

equipped with a 66 m.times.2.6.

DETD X-ray crystallography has been used to elucidate the molecular geometry of the trisaccharide triterpene asiaticoside from Centella

asiatica (Umbelliferae). Crystallization was from dioxane

(Mahato et al., 1987). X-ray diffraction analysis was also successful

for confirmation of the structure.

DETD . . . end of the study. Groups 2-4 are fed the NRC diet plus 1% cholesterol for another two-week period to induce hypercholesterolemia. Then, Group 2 will continue on this diet until the end of the study, while Groups 3 and 4 will. . .

=> s weight loss or obesi? or weight gain? L22 321718 WEIGHT LOSS OR OBESI? OR WEIGHT GAIN?

=> s 16 and 122

L23 15 L6 AND L22

=> dup rem 123

PROCESSING COMPLETED FOR L23

L24 13 DUP REM L23 (2 DUPLICATES REMOVED)

=> d ibib abs 9-13

L24 ANSWER 9 OF 13 MEDLINE DUPLICATE 1

ACCESSION NUMBER: 2001332231

001332231 MEDLINE

DOCUMENT NUMBER: 21293407 PubMed ID: 11399290

TITLE: Protection against radiation-induced conditioned taste

aversion by Centella asiatica.

AUTHOR:

Shobi V; Goel H C

CORPORATE SOURCE: Radiation Biology Division, Institute of Nuclear Medicine

and Allied Sciences, DRDO, Lucknow Marg, New Delhi 110 054,

India.

SOURCE: PHYSIOLOGY AND BEHAVIOR, (2001 May) 73 (1-2) 19-23.

Journal code: 0151504. ISSN: 0031-9384.

PUB. COUNTRY: United States

DOCUMENT TYPE: Journal; Article; (JOURNAL ARTICLE)

LANGUAGE: English

FILE SEGMENT: Priority Journals

ENTRY MONTH: 200108

ENTRY DATE: Entered STN: 20010813

Last Updated on STN: 20010813 Entered Medline: 20010809

AB Radiations are known to cause behavioural perturbations like conditioned taste aversion (CTA), performance decrement, learning, etc., even at very low doses. The manifestation of radiation-induced behavioural degradation has not been understood well and requires further studies. Therefore, the effects of low-dose whole-body 60Co gamma-irradiation in male rats were

studied in terms of body weight and CTA learning. For CTA, the consumption of saccharin solution was considered as a parameter. To protect against the adverse effects of radiation, Centella asiatica (aqueous extract) was tested and compared with ondansetron, a standard antiemetic drug. A dose of 2 Gy incurred significant body weight loss [t(9)=9.00, P<.05] and induced CTA in rats [t(26)=9.344,P<.01]. Administration of C. asiatica (100 mg/kg bw ip, 2 Gy, -1 h) rendered significant radioprotection against radiation-induced body weight loss and CTA that became evident on the second postirradiation day [t(7)=0.917, P>>.05; t(7)=4.016, P>.05]. Ondansetron (1 mg/kg bw) elicited higher degree of protection against CTA [t(7)=3.641, P>.05] than C. asiatica [t(7)=7.196, P>.05] on the first postirradiation day, but on the second postirradiation day, both were equally effective [t(7)=3.38, P>.05; t(7)=4.01, P>.05]. In case of C. asiatica-treated animals, however, there was a consistently declining CTA from the second to the fifth postirradiation day whereas in ondansetron-treated animals it was inconsistent. Present investigation suggests that C. asiatica could be useful in preventing radiation-induced behavioural changes during clinical radiotherapy.

L24 ANSWER 10 OF 13 USPATFULL

ACCESSION NUMBER: 93:20355 USPATFULL

TITLE: Slimming composition based on Ginkgo biloba as an

alpha-2-blocker

INVENTOR(S): Soudant, Etienne, Fresnes, France

Nadaud, Jean-Francois, Paris, France

PATENT ASSIGNEE(S): L'Oreal, Paris, France (non-U.S. corporation)

> NUMBER KIND DATE -----

PATENT INFORMATION: US 5194259 APPLICATION INFO.: US 1991-798329 19911127 (7)

> NUMBER DATE -----

PRIORITY INFORMATION: FR 1990-14864 19901128

DOCUMENT TYPE: Utility Granted FILE SEGMENT:

PRIMARY EXAMINER: Page, Thurman K. ASSISTANT EXAMINER: Hulina, Amy

LEGAL REPRESENTATIVE: Cushman, Darby & Cushman

NUMBER OF CLAIMS: 10 EXEMPLARY CLAIM: 1 LINE COUNT: 382

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

AB A cosmetic slimming composition for topical application to the skin contains in combination Ginkgo biloba as an alpha-2-blocker and at least one other alpha-2-blocker. This anti-cellulitis composition is capable of checking or stopping local fat accumulation and improving the esthetic appearance of the skin.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

L24 ANSWER 11 OF 13 BIOSIS COPYRIGHT 2003 BIOLOGICAL ABSTRACTS INC.

ACCESSION NUMBER: 1993:413231 BIOSIS DOCUMENT NUMBER: PREV199396078956

Plants used in ethnomedicine for asthma in Kivu (Zaire. TITLE: AUTHOR(S): Kasonia, K. (1); Ansay, M.; Gustin, P.; Plume, C.

CORPORATE SOURCE: (1) Universite Lubumbashi, l'Universite Liege, Fac.

Medecine Veterinaire, Pharmacologie Toxicologie, B-41 Bld.

de Colonster, Sart-Tilman, B-4000 Liege, Belgique Belgian Journal of Botany, (1993) Vol. 126, No. 1, pp.

20-28.

ISSN: 0778-4031.

DOCUMENT TYPE: Article LANGUAGE: French

SOURCE:

SUMMARY LANGUAGE: French; English

In Kivu (Zaire), out ethnobotanical investigations led to the identification of 30 plants belonging to 20 families used in the treatment of asthma. They are listed with scientific names, but local names, preparation methods and direction for use are also detailed.

92:96754 USPATFULL ACCESSION NUMBER:

TITLE: Magnetically influenced homeopathic pharmaceutical

formulations, methods of their preparation and methods

of their administration

INVENTOR(S): Whitson-Fischman, Walter, New York, NY, United States PATENT ASSIGNEE(S):

Whitson Laboratories, Inc., New York, NY, United States

(U.S. corporation)

NUMBER KIND DATE PATENT INFORMATION: US 5162037 19921110 APPLICATION INFO.: US 1991-696759 19910507 (7)

RELATED APPLN. INFO.: Continuation-in-part of Ser. No. US 1990-540295, filed on 19 Jun 1990, now abandoned which is a continuation

of Ser. No. US 1988-176731, filed on 1 Apr 1988, now

abandoned Utility

DOCUMENT TYPE: FILE SEGMENT: Granted PRIMARY EXAMINER: Cohen, Lee S.

LEGAL REPRESENTATIVE: Haug, Edgar H., Kilcoyne, John M.

NUMBER OF CLAIMS: 38

EXEMPLARY CLAIM:

NUMBER OF DRAWINGS: 4 Drawing Figure(s); 3 Drawing Page(s)

LINE COUNT: 2156

A method for treating pathogenic conditions of the human body by preparing a homeopathic mixture of at least one herb, herbal extract or other compound exhibiting therapeutic properties, adding a magnetically permeable substance to the mixture if necessary, magnetizing the resulting mixture to impart a substantially unipolar magnetic charge on the mixture and administering the magnetized mixture through one or more specific acupuncture points associated with producing a desired response to the particular condition being treated. The invention is also directed to the treatment of various diseases through the oral, auricular, topical or injectable administration of magnetically influenced homeopathic medicaments.

L24 ANSWER 13 OF 13 BIOSIS COPYRIGHT 2003 BIOLOGICAL ABSTRACTS INC.

ACCESSION NUMBER: 1993:413230 BIOSIS DOCUMENT NUMBER: PREV199396078955

TITLE: Evaluation of possible galactagogue activity of a selected

group of Sri Lankan medicinal plants.

AUTHOR (S): Tennekoon, Kamani H. (1); Jeevathayaparan, S.;

Karunanayake, Eric H.

CORPORATE SOURCE: (1) Dep. Physiol., Fac. Med., Univ. Colombo, Colombo SOURCE:

Journal of the National Science Council of Sri Lanka,

(1992) Vol. 20, No. 1, pp. 33-41.

ISSN: 0300-9254.

DOCUMENT TYPE: Article LANGUAGE: English

In view of the potential benefits of herbal medicinal plants in improving lactation, a selected group of Sri Lankan plants used in the traditional system of Medicine as galactagogues were evaluated for possible galactagoguic activity of Sprague Dawley rats. Extracts of Nigella sativa, Dregea volubilis, Ipomea digitata, Borassus flabellifer, Corriandrum sativum, Momordica charantia, Carica papaya, Centella asiatica and Allium sativum were administered orally to lactating rates for 1 week from the 5th day after delivery. The number of littermates were limiting to 7 per mother. Weight gain and milk intake in the litter on the 7th day of the experiment were compared with control groups that received equivalent amounts of vehicle under identical conditions. Significant galactagoguic activity was not detected in any of the plant extracts studied, although the doses of extracts used were adequate to concentrate possible active principles. However, a seasonal variation of the active principle or possible inactivation of the galatagoguic activity in the digestive system cannot be excluded.

=> s urinat?

18178 URINAT? L25

```
L26 1 L6 AND L25
```

=> d

L26 ANSWER 1 OF 1 USPATFULL AN 1999:91815 USPATFULL

TI Haemostatic circumcision bandage

IN Friedman, Jack, 5050 Bourret, #209, Montreal Que., Canada H3W1L4

PI US 5935091 19990810 AI US 1997-993095 19971218 (8)

DT Utility FS Granted

LN.CNT 231

INCL INCLM: 602/079.000

INCLS: 602/041.000; 602/058.000; 602/067.000; 128/844.000; 002/021.000;

604/037.000

NCL NCLM: 602/079.000

NCLS: 002/021.000; 128/844.000; 602/041.000; 602/058.000; 602/067.000;

604/037.000

IC [6]

ICM: A61F013-00

EXF 602/41; 602/79; 604/37

=> s madasiatic acid or asiaticoside

L27 390 MADASIATIC ACID OR ASIATICOSIDE

=> s 127 and (17 or diuret?)

L28 9 L27 AND (L7 OR DIURET?)

=> dup rem 128

PROCESSING COMPLETED FOR L28

L29 9 DUP REM L28 (0 DUPLICATES REMOVED)

=> d ibib abs kwic 5-9

PATENT INFORMATION:

L29 ANSWER 5 OF 9 USPATFULL

ACCESSION NUMBER: 2001:59406 USPATFULL

TITLE: Solubility parameter based drug delivery system and

method for altering drug saturation concentration

INVENTOR(S): Miranda, Jesus, Miami, FL, United States

Sablotsky, Steven, Miami, FL, United States

PATENT ASSIGNEE(S): Noven Pharmaceuticals, Inc., Miami, FL, United States

(U.S. corporation)

APPLICATION INFO.: US 1999-318121 19990525 (9)
RELATED APPLN. INFO.: Division of Ser. No. US 1997-907906, filed on 11 Aug

1997 Continuation-in-part of Ser. No. US 1994-178558, filed on 7 Jan 1994, now patented, Pat. No. US 5656286,

issued on 12 Aug 1997

DOCUMENT TYPE: Utility

FILE SEGMENT: Granted

PRIMARY EXAMINER: Dodson, Shelley A.
ASSISTANT EXAMINER: Williamson, Michael A.

LEGAL REPRESENTATIVE: Foley & Lardner

NUMBER OF CLAIMS: 4 EXEMPLARY CLAIM: 1

NUMBER OF DRAWINGS: 20 Drawing Figure(s); 19 Drawing Page(s)

LINE COUNT: 3035

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

AB A blend of at least two polymers, or at least one polymer and a soluble polyvinylpyrrolidone, in combination with a drug provides a pressure-sensitive adhesive composition for a transdermal drug delivery system in which the drug is delivered from the pressure-sensitive adhesive composition and through dermis when the pressure-sensitive adhesive composition is in contact with human skin. According to the invention, soluble polyvinylpyrrolidone can be used to prevent crystallization of the drug, without affecting the rate of drug delivery from the pressure-sensitive adhesive composition.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

92. Diuretics, including: DETD

143. Vulnerary agents such as Acetylcysteine, Allantoin, DETD Asiaticoside, Cadexomer Iodine, Chitin, Dextranomer and Oxaceprol.

L29 ANSWER 6 OF 9 USPATFULL

ACCESSION NUMBER:

2000:18064 USPATFULL

TITLE:

Solubility parameter based drug delivery system and method for altering drug saturation concentration

INVENTOR (S):

Miranda, Jesus, Miami, FL, United States Sablotsky, Steven, Miami, FL, United States

PATENT ASSIGNEE(S):

Noven Pharmaceuticals, Inc., Miami, FL, United States

(U.S. corporation)

NUMBER KIND DATE -----PATENT INFORMATION: US 6024976 20000215

APPLICATION INFO. : US 1997-907906 19970811 (8) RELATED APPLN. INFO.: Continuation of Ser. No. US 1994-178558, filed on 7 Jan

1994, now patented, Pat. No. US 5656286 which is a continuation-in-part of Ser. No. US 1991-722342, filed on 27 Jun 1991 which is a continuation-in-part of Ser.

No. US 671709

DOCUMENT TYPE: FILE SEGMENT:

Utility Granted

PRIMARY EXAMINER: LEGAL REPRESENTATIVE: Foley & Lardner

Venkat, Jyothsna

NUMBER OF CLAIMS: EXEMPLARY CLAIM:

1

NUMBER OF DRAWINGS:

20 Drawing Figure(s); 19 Drawing Page(s)

LINE COUNT:

3328

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

A blend of at least two polymers, or at least one polymer and a soluble polyvinylpyrrolidone, in combination with a drug provides a pressure-sensitive adhesive composition for a transdermal drug delivery system in which the drug is delivered from the pressure-sensitive adhesive composition and through dermis when the pressure-sensitive adhesive composition is in contact with human skin. According to the invention, soluble polyvinylpyrrolidone can be used to prevent crystallization of the drug, without affecting the rate of drug delivery from the pressure-sensitive adhesive composition.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

DETD 92. Diuretics, including:

DETD 143. Vulnerary agents such as Acetylcysteine, Allantoin, Asiaticoside, Cadexomer Iodine, Chitin, Dextranomer and Oxaceprol.

L29 ANSWER 7 OF 9 USPATFULL

ACCESSION NUMBER:

1998:17360 USPATFULL

TITLE:

Compositions and methods for topical administration of

pharmaceutically active agents

INVENTOR (S):

Kanios, David P., Miami, FL, United States Gentile, Joseph A., Plantation, FL, United States Mantelle, Juan A., Miami, FL, United States Sablotsky, Steven, Miami, FL, United States

PATENT ASSIGNEE(S):

Noven Pharmaceuticals, Inc., Miami, FL, United States

(U.S. corporation)

NUMBER KIND DATE -----US 5719197 19980217 US 1995-477361 19950607 (8)

PATENT INFORMATION: APPLICATION INFO.: RELATED APPLN. INFO.:

Continuation-in-part of Ser. No. US 1993-112330, filed on 27 Aug 1993, now patented, Pat. No. US 5446070 which is a continuation-in-part of Ser. No. US 1991-813196, filed on 23 Dec 1991, now patented, Pat. No. US 5234957 which is a continuation-in-part of Ser. No. US 1991-661827, filed on 27 Feb 1991, now abandoned , said Ser. No. US 1995-477361, filed on 7 Jun 1995 which is a continuation-in-part of Ser. No. US 1993-67001, filed on 26 May 1993 which is a continuation of Ser. No. US

1991-671709, filed on 2 Apr 1991, now patented, Pat. No. US 5300291 which is a continuation-in-part of Ser. No. US 1989-295847, filed on 11 Jan 1989, now patented, Pat. No. US 4994267 which is a continuation-in-part of Ser. No. US 1988-164482, filed on 4 Mar 1988, now

patented, Pat. No. US 4814168

DOCUMENT TYPE: Utility FILE SEGMENT: Granted

PRIMARY EXAMINER: Azpuru, Carlos A. LEGAL REPRESENTATIVE: Foley & Lardner

NUMBER OF CLAIMS: 27
EXEMPLARY CLAIM: 1
LINE COUNT: 1799

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

AB Compositions for topical application comprising a therapeutically effective amount of a pharmaceutical agent(s), a pharmaceutically acceptable bioadhesive carrier, a solvent for the pharmaceutical agent(s) in the carrier and a clay, and methods of administering the pharmaceutical agents to a mammal are disclosed.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

DETD DIURETIC

DETD VULNERARY such as Allantoin, Asiaticoside, Cadexomer Iodine, Chitin, Dextranomer

L29 ANSWER 8 OF 9 USPATFULL

ACCESSION NUMBER: 97:70731 USPATFULL

TITLE: Solubility parameter based drug delivery system and method for altering drug saturation concentration

INVENTOR(S): Miranda, Jesus, Miami, FL, United States

Sablotsky, Steven, Miami, FL, United States

PATENT ASSIGNEE(S): Noven Pharmaceuticals, Inc., Miami, FL, United States

(U.S. corporation)

 PATENT INFORMATION:
 US 5656286
 19970812

 APPLICATION INFO.:
 US 1994-178558
 19940107 (8)

 RELATED APPLN. INFO.:
 Continuation-in-part of Ser. No. US 1991-722342, filed

on 27 Jun 1991, now patented, Pat. No. US 5474783 which is a continuation-in-part of Ser. No. US 1991-671709, filed on 2 Apr 1991, now patented, Pat. No. US 5300291 which is a continuation-in-part of Ser. No. US 1989-295847, filed on 11 Jan 1989, now patented, Pat. No. US 4994267, issued on 19 Feb 1991 which is a

continuation-in-part of Ser. No. US 1988-164482, filed on 4 Mar 1988, now patented, Pat. No. US 4814168,

issued on 21 Mar 1989

DOCUMENT TYPE: Utility FILE SEGMENT: Granted

PRIMARY EXAMINER: Venkat, Jyothsna LEGAL REPRESENTATIVE: Foley & Lardner NUMBER OF CLAIMS: 73

EXEMPLARY CLAIM: 1,4

NUMBER OF DRAWINGS: 20 Drawing Figure(s); 19 Drawing Page(s)

LINE COUNT: 3344

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

AB A blend of at least two polymers, or at least one polymer and a soluble polyvinylpyrrolidone, in combination with a drug provides a pressure-sensitive adhesive composition for a transdermal drug delivery system in which the drug is delivered from the pressure-sensitive adhesive composition and through dermis when the pressure-sensitive adhesive composition is in contact with human skin. According to the invention, soluble polyvinylpyrrolidone can be used to prevent crystallization of the drug, without affecting the rate of drug delivery from the pressure-sensitive adhesive composition.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

DETD 92. Diuretics, including:

DETD 143. Vulnerary agents such as Acetylcysteine, Allantoin, Asiaticoside, Cadexomer Iodine, Chitin, Dextranomer and Oxaceprol.

L29 ANSWER 9 OF 9 USPATFULL

ACCESSION NUMBER: 95:78209 USPATFULL

TITLE: Compositions and methods for topical administration of

pharmaceutically active agents

INVENTOR(S): Mantelle, Juan A., Miami, FL, United States

PATENT ASSIGNEE(S): Nover Pharmaceuticals, Inc., Miami, FL, United States

(U.S. corporation)

DISCLAIMER DATE: 20100810

RELATED APPLN. INFO.: Continuation-in-part of Ser. No. US 1991-813196, filed on 23 Dec 1991, now patented, Pat. No. US 5234957 which

is a continuation-in-part of Ser. No. US 1991-661827,

filed on 27 Feb 1991, now abandoned

DOCUMENT TYPE: Utility FILE SEGMENT: Granted

PRIMARY EXAMINER: Page, Thurman K.
ASSISTANT EXAMINER: Azpuru, Carlos
LEGAL REPRESENTATIVE: Foley & Lardner

NUMBER OF CLAIMS: 45 EXEMPLARY CLAIM: 1 LINE COUNT: 24

2434

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

AB Compositions for topical application comprising a therapeutically effective amount of a pharmaceutical agent(s), a pharmaceutically acceptable carrier, and a solvent for the pharmaceutical agent(s) in the carrier and methods of administering the pharmaceutical agents to a mammal are disclosed.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

DETD DIURETIC

DETD VULNERARY such as Allantoin, Asiaticoside, Cadexomer Iodine, Chitin, Dextranomer

=> d ti tot

- L29 ANSWER 1 OF 9 USPATFULL
- TI Compositions and methods to effect the release profile in the transdermal administration of active agents
- L29 ANSWER 2 OF 9 USPATFULL
- TI Triterpene compositions and methods for use thereof
- L29 ANSWER 3 OF 9 USPATFULL
- TI Cosmetic formulations containing extracts from phyllanthus emblica and centella asiatica and/or bacopa monnieri
- L29 ANSWER 4 OF 9 USPATFULL
- TI Cosmetic preparations containing extracts from phyllanthus emblica and centella asiatica and/or bacopa monnieri
- L29 ANSWER 5 OF 9 USPATFULL
- TI Solubility parameter based drug delivery system and method for altering drug saturation concentration
- L29 ANSWER 6 OF 9 USPATFULL
- TI Solubility parameter based drug delivery system and method for altering drug saturation concentration
- L29 ANSWER 7 OF 9 USPATFULL
- TI Compositions and methods for topical administration of pharmaceutically active agents
- L29 ANSWER 8 OF 9 USPATFULL
- TI Solubility parameter based drug delivery system and method for altering drug saturation concentration
- L29 ANSWER 9 OF 9 USPATFULL
- TI Compositions and methods for topical administration of pharmaceutically

=> d ibib abs kwic 2 3

L29 ANSWER 2 OF 9 USPATFULL

2002:224280 USPATFULL ACCESSION NUMBER:

TITLE: Triterpene compositions and methods for use thereof

Arntzen, Charles J., Ithaca, NY, United States INVENTOR (S):

Blake, Mary E., Tucson, AZ, United States Gutterman, Jordan U., Houston, TX, United States Hoffmann, Joseph J., Tucson, AZ, United States

Jayatilake, Gamini S., Broomfield, CO, United States Bailey, David T., Boulder, CO, United States

PATENT ASSIGNEE(S): Research Development Foundation, Carson City, NV,

United States (U.S. corporation)

KIND DATE NUMBER ••••• PATENT INFORMATION: US 6444233 B1 20020903 APPLICATION INFO.: US 1999-314691 19990519 (9)

> NUMBER DATE

US 1998-99066P US 1998-85997P PRIORITY INFORMATION: 19980903 (60)

19980519 (60)

DOCUMENT TYPE: Utility FILE SEGMENT: GRANTED

PRIMARY EXAMINER: Tate, Christopher R. ASSISTANT EXAMINER: Flood, Michele C. LEGAL REPRESENTATIVE: Fulbright & Jaworski LLP

NUMBER OF CLAIMS: 18 EXEMPLARY CLAIM:

NUMBER OF DRAWINGS: 73 Drawing Figure(s); 43 Drawing Page(s)

LINE COUNT: 7526

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

The invention provides novel saponin mixtures and compounds which are AB isolated from the species Acacia victoriae and methods for their use. These compounds may contain a triterpene moiety, such as acacic or oleanolic acid, to which oligosaccharides and monoterpenoid moieties are attached. The mixtures and compounds have properties related to the regulation of apoptosis and cytotoxicity of cells and exhibit potent anti-tumor effects against a variety of tumor cells.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

. . . invention may be used as solvents, antioxidants, anti-fungal SUMM and anti-viral agents, piscicides or molluscicides, contraceptives, antihelmintics, angiogenesis regulators, UV-protectants, expectorants, diuretics, anti-inflammatory agents, regulators of cholesterol metabolism, cardiovascular effectors, anti-ulcer agents, analgesics, sedatives, immunomodulators, antipyretics, as agents for decreasing capillary fragility,.

DETD The two main saponins asiaticoside and madecassoside from Centella asiatica (Umbelliferae) have been separated with the aid of an Ito multi-layer coil separator-extractor (P.C. Inc.).

DETD X-ray crystallography has been used to elucidate the molecular geometry of the trisaccharide triterpene asiaticoside from Centella asiatica (Umbelliferae). Crystallization was from dioxane (Mahato et al., 1987). X-ray diffraction analysis was also successful for confirmation.

DETD . of the triterpene compounds of the invention as solvents, anti-fungal and anti-viral agents, piscicides or molluscicides, contraceptives, antihelmintics, UV-protectants, expectorants, diuretics, anti-inflammatory agents, regulators of cholesterol metabolism, cardiovascular effectors, anti-ulcer agents, analgesics, sedatives, immunomodulators, antipyretics, angiogenesis regulators, as agents for decreasing. .

DETD . the use of the compounds of the invention as anti-fungal and anti-viral agents, piscicides or molluscicides, contaceptives, anthelmintics, UV-protectants, expectorants, diuretics, anti-inflammatory agents, regulators of cholesterol metabolism, cardiovascular effectors, anti-ulcer agents, analgesics, sedatives, immunomodulators, antipyretics, regulators of angiogenesis, and as

agents. . .

L29 ANSWER 3 OF 9 USPATFULL

ACCESSION NUMBER: 2001:139171 USPATFULL

TITLE: Cosmetic formulations containing extracts from

phyllanthus emblica and centella asiatica and/or bacopa

monnieri

INVENTOR(S): Singh-Verma, Shyam B., Kerpen, Germany, Federal

Republic of

RELATED APPLN. INFO.: Continuation of Ser. No. US 1999-331791, filed on 27

Jul 1999, PENDING

NUMBER DATE

PRIORITY INFORMATION: DE 1996-19654635 19961228

DOCUMENT TYPE: Utility FILE SEGMENT: APPLICATION

LEGAL REPRESENTATIVE: BANNER & WITCOFF, 1001 G STREET N W, SUITE 1100,

WASHINGTON, DC, 20001

NUMBER OF CLAIMS: 7
EXEMPLARY CLAIM: 1
LINE COUNT: 482

AB The present invention relates to cosmetic formulations for topical application containing extracts from Phyllanthus emblica and Centella asiatica and/or Bacopa monnieri, and the use of such formulations for

the care of the human skin.

In particular, the present invention relates to cosmetic formulations for topical application containing extracts from Phyllanthus emblica and Centella asiatica and/or Bacopa monnieri in addition to per se known adjuvants and expedients.

SUMM . . . primary tincture prepared according to HRB I, there are mentioned the alcaloid hydrocotyline, the triterpenic acids (asiatic acid, madecassic acid, madasiatic acid) and the triterpene saponin asiaticoside (Haager's Handbuch der Drogenkunde). The oral application of an infusion of the medicinal plant is said to have blood-purifying, tonicising and diuretic properties. When applied topically, the extracts and tinctures have antiphlogistic, antibacterial and wound-healing effects which are said

to be attributable. . .

SUMM . . . Springer Verlag, 1972, 3rd volume, p. 792-793, it is known that the active substance madecassoid has an anti-inflammatory effect, while asiaticoside, which stimulates mitoses, promotes the healing of premitis and wounds.

=> d ibib 4

L29 ANSWER 4 OF 9 USPATFULL

ACCESSION NUMBER: 2001:111876 USPATFULL

TITLE: Cosmetic preparations containing extracts from

phyllanthus emblica and centella asiatica and/or bacopa

19990727 PCT 102(e) date

monnieri

INVENTOR(S): Singh-Verma, Shyam B., Nubhaumallee 13, 50169, Kerpen,

Germany, Federal Republic of

KIND DATE NUMBER ----------PATENT INFORMATION: US 6261605 B1 20010717 WO 9829089 19980709 APPLICATION INFO.: US 1999-331791 19990727 (9) WO 1997-EP7113 19971218 19990727 PCT 371 date

NUMBER DATE

PRIORITY INFORMATION: DE 1996-19654635 19961228

DOCUMENT TYPE: Utility FILE SEGMENT: GRANTED

PRIMARY EXAMINER: Prats, Francisco
ASSISTANT EXAMINER: Coe, Susan D.
LEGAL REPRESENTATIVE: Banner & Witcoff, Ltd.

NUMBER OF CLAIMS: 8 EXEMPLARY CLAIM: 1 LINE COUNT: 487

=>

=> fil stng

COST IN U.S. DOLLARS SINCE FILE TOTAL ENTRY SESSION FULL ESTIMATED COST 103.21 151.74

DISCOUNT AMOUNTS (FOR QUALIFYING ACCOUNTS)
SINCE FILE TOTAL
ENTRY SESSION
CA SUBSCRIBER PRICE
-1.95
-2.60

FILE 'STNGUIDE' ENTERED AT 11:16:00 ON 08 JAN 2003
USE IS SUBJECT TO THE TERMS OF YOUR CUSTOMER AGREEMENT
COPYRIGHT (C) 2003 AMERICAN CHEMICAL SOCIETY, JAPAN SCIENCE
AND TECHNOLOGY CORPORATION, AND FACHINFORMATIONSZENTRUM KARLSRUHE

FILE CONTAINS CURRENT INFORMATION.
LAST RELOADED: Dec 20, 2002 (20021220/UP).

=> FIL MEDL CAPL BIOSIS USPATF

COST IN U.S. DOLLARS SINCE FILE TOTAL ENTRY SESSION FULL ESTIMATED COST 0.36 152.10

DISCOUNT AMOUNTS (FOR QUALIFYING ACCOUNTS)

SINCE FILE
ENTRY
SESSION
CA SUBSCRIBER PRICE

0.00
-2.60

FILE 'MEDLINE' ENTERED AT 11:19:49 ON 08 JAN 2003

FILE 'CAPLUS' ENTERED AT 11:19:49 ON 08 JAN 2003 USE IS SUBJECT TO THE TERMS OF YOUR STN CUSTOMER AGREEMENT. PLEASE SEE "HELP USAGETERMS" FOR DETAILS. COPYRIGHT (C) 2003 AMERICAN CHEMICAL SOCIETY (ACS)

FILE 'BIOSIS' ENTERED AT 11:19:49 ON 08 JAN 2003 COPYRIGHT (C) 2003 BIOLOGICAL ABSTRACTS INC.(R)

FILE 'USPATFULL' ENTERED AT 11:19:49 ON 08 JAN 2003
CA INDEXING COPYRIGHT (C) 2003 AMERICAN CHEMICAL SOCIETY (ACS)

=> s selenium

L30 128947 SELENIUM

=> s 130 and 111

L31 207 L30 AND L11

=> s 130 (S) 111

L32 49 L30 (S) L11

=> dup rem 132

PROCESSING COMPLETED FOR L32

L33 35 DUP REM L32 (14 DUPLICATES REMOVED)

=> d ibib abs kwic 33-35

L33 ANSWER 33 OF 35 MEDLINE

ACCESSION NUMBER: 87260478 MEDLINE

DOCUMENT NUMBER: 87260478 PubMed ID: 3037508

TITLE: Key issues in nutrition. Disease prevention through

adulthood and old age.

AUTHOR: Fahey P J; Boltri J M; Monk J S

SOURCE: POSTGRADUATE MEDICINE, (1987 Jul) 82 (1) 135-42.

Journal code: 0401147. ISSN: 0032-5481.

PUB. COUNTRY: United States

DOCUMENT TYPE: Journal; Article; (JOURNAL ARTICLE)

LANGUAGE: English

FILE SEGMENT: Abridged Index Medicus Journals; Priority Journals

ENTRY MONTH: 198708

ENTRY DATE: Entered STN: 19900305

Last Updated on STN: 19900305 Entered Medline: 19870804

Certain dietary practices are valid methods of lowering the risk of AB disease. Others, while popular, have unproven benefits or may even be associated with risks of their own. Careful evaluation of hypercholesterolemia is necessary. Persons with a high level of low-density lipoprotein (LDL) cholesterol and a low level of high-density lipoprotein (HDL) cholesterol need diet therapy, because they are at increased risk of cardiovascular disease. Weight reduction and fat restriction can lower blood pressure, help control hyperglycemia, and improve the LDL cholesterol-HDL cholesterol ratio. Some evidence indicates a protective role of beta carotene against cancer in animals. However, hypervitaminosis A is dangerous and relatively easy to accomplish, so supplementation beyond a multivitamin tablet is discouraged. Data about inhibition of cancer in humans through use of high doses of vitamin E or C or selenium are inconclusive, and studies of effects of long-term ingestion are not available. In general, megadoses of even healthy substances are thought to be dangerous. Decreased consumption of fat, increased consumption of foods high in fiber, and elimination of alcohol and tobacco are sensible recommendations. Consumption of cruciferous vegetables has not been proven to reduce the incidence of cancer, but a moderate amount of them in the diet would seem reasonable. AB . . . disease. Others, while popular, have unproven benefits or may even be associated with risks of their own. Careful evaluation of hypercholesterolemia is necessary. Persons with a high level of low-density lipoprotein (LDL) cholesterol and a low level of high-density lipoprotein (HDL). . . is discouraged. Data about inhibition of cancer in humans through use of high doses of vitamin E or C or selenium are inconclusive, and studies of effects of long-term ingestion are not

L33 ANSWER 34 OF 35 BIOSIS COPYRIGHT 2003 BIOLOGICAL ABSTRACTS INC.

ACCESSION NUMBER: 1985:412088 BIOSIS

DOCUMENT NUMBER: BA80:82080

TITLE: RELATIONSHIPS BETWEEN TRACE ELEMENTS AND ATHEROSCLEROSIS.

available. In general, megadoses of even healthy substances are. . .

AUTHOR(S): AALBERS T G; HOUTMAN J P W

CORPORATE SOURCE: RIVM, P.O. BOX 1, 3720 BA BILTHOVEN, NETHERLANDS.

SOURCE: SCI TOTAL ENVIRON, (1985) 43 (3), 255-284.

CODEN: STEVA8. ISSN: 0048-9697.

FILE SEGMENT: BA; OLD LANGUAGE: English

The possible relationship between trace element (Al, As, Cd, Co, Cr, Cu, Fe, Hg, Mn, Mo, Ni, Pb, Sb, Se, Zn) concentrations in various human tissues (heart, liver, kidney, aorta, rib and head hair) and cardiovascular diseases was studied on the basis of indications in the literature that trace elements may be directly or indirectly involved in cardiovascular disease processes. The underlying theme was that (slightly) reduced, as well as (slightly) elevated, concentrations compared with optimum values could, in the long term, lead to atherosclerotic lesions. In this project the tissues were obtained by autopsy involving 200 individuals (hospitalized patients and victims of traffic accidents). The seriousness of cardiovascular disease was quantitatively expressed by the degree of atherosclerosis of the descending branch of the left coronary artery (LAD) and of the abdominal aorta, for which a special measurement method was developed. Correlations were evaluated by 2 different methods, i.e., by a comparison of patients with extremely high or extremely low degrees of atherosclerosis and by means of stepwise multiple linear regression (SMLR) analysis. Corrections were made for the influence of age. The element Cd was found to be positively, and the elements Cu, Co, Se and Zn negatively, correlated with the degree of atherosclerosis. The inclusion of risk factors (diabetes mellitus, hypercholesterolemia, hypertension, obesity and smoking) did not improve the explained variance. тт

IT Miscellaneous Descriptors

DISEASE OBESITY CHROMIUM ARSENIC CADMIUM ALUMINUM COBALT SELENIUM ZINC COPPER IRON MERCURY MANGANESE MOLYBDENUM NICKEL LEAD ANTIMONY

L33 ANSWER 35 OF 35 CAPLUS COPYRIGHT 2003 ACS ACCESSION NUMBER: 1984:417136 CAPLUS

DOCUMENT NUMBER: 101:17136

TTTLE Action of bioflavonoids on lipid peroxidation and

glutathione redox system in hypercholesterolemic rats

Rathi, Ashok B.; Nath, N.; Chari, S. N. AUTHOR (S): CORPORATE SOURCE: Dep. Biochem., Nagpur Univ., Nagpur, India

SOURCE:

Indian Journal of Medical Research (1913-1988) (1984),

79(April), 508-13 CODEN: IJMRAO; ISSN: 0019-5340

DOCUMENT TYPE: Journal LANGUAGE: English

Enhanced hepatic lipid peroxidn., decreased activity of the glutathione redox system and Se depletion were obsd. in cholesterol-cholic acid induced hypercholesterolemic rats. These aberrations were significantly reversed by simultaneous administration of quercetin [117-39-5] or hesperidin [520-26-3]. The action of these bioflavonoids on the lipoperoxide formation is not certain. However, it is possible that these compds. restore the content of glutathione and the activity of glutathione peroxidase [9013-66-5] by retarding the depletion of Se in the hypercholesterolemic condition. The mechanism of action of these compds. is discussed.

selenium liver hypercholesterolemia bioflavonoid; bioflavonoid lipid peroxidn hypercholesterolemia; glutathione redox system hypercholesterolemia bioflavonoid

=> d ibib abs kwic 20-23

L33 ANSWER 20 OF 35 MEDLINE

ACCESSION NUMBER: 2000220279 MEDLINE

DOCUMENT NUMBER: 20220279 PubMed ID: 10756780

TITLE: [The importance of the use of selenium in the role of an antioxidant in preventing cardiovascular diseases].

Importanta utilizarii seleniului cu rol antioxidant in

preventia bolilor cardiovasculare.

AUTHOR: Azoicai D; Ivan A; Bradatean M; Pavel M; Jerca L;

Iacobovici A; Popovici I; Gheorghita N

Disciplina de Epidemiologie, Facultatea de Medicina, CORPORATE SOURCE:

Universitatea de Medicina si Farmacie Gr. T. Popa, Iasi. SOURCE: REVISTA MEDICO-CHIRURGICALA A SOCIETATII DE MEDICI SI

NATURALISTI DIN IASI, (1997 Jul-Dec) 101 (3-4) 109-15.

Journal code: 0413735. ISSN: 0300-8738.

PUB. COUNTRY: Romania

DOCUMENT TYPE: Journal; Article; (JOURNAL ARTICLE)

LANGUAGE: Romanian

FILE SEGMENT: Priority Journals

ENTRY MONTH: 200005

ENTRY DATE: Entered STN: 20000606

Last Updated on STN: 20000606

Entered Medline: 20000519

AB The evaluation of the results of the oxygen free radicals (RLO2) formation is a current subject in biology and medicine. The oxidative stress, which is the altering of the balance between the higher activity of oxygen and the enzymatic or nonenzymatic protection systems, may be one of the causes that starts and aggravates a disease. In this context, the supplementation of the diet with selenium, superoxide dismutase, vitamins A, C, E, is considered a primary prevention measure (for the apparently healthy persons) and a secondary one (for those with advancing forms of disease) that is both efficient and modern by utilization of some "drug-food" products. The transversal study realized on a group of 39 blood donors presence of the cardiovascular risk determined by the raising of the prevalence of some atherogenic factors (active smoking, hypercholesterolemia) which is also expressed by the lowering of the level of some oxidative stress indicators (glutathione peroxidase--GSH-Px < 0.139 moli/ml and catalase < 2.20 U/ml). The simultaneous low intake of selenium from the central drinking water supplies in the city of Iasi (0.1-1 g/l) has permitted us to consider necessary the diet supplementation both with foods rich in

vitamins with an antioxidant role and with specific medication with selenium, as a protective micro-element.

AB . . be one of the causes that starts and aggravates a disease. In this context, the supplementation of the diet with selenium, superoxide dismutase, vitamins A, C, E, is considered a primary prevention measure (for the apparently healthy persons) and a secondary. . . blood donors presence of the cardiovascular risk determined by the raising of the prevalence of some atherogenic factors (active smoking, hypercholesterolemia) which is also expressed by the lowering of the level of some oxidative stress indicators (glutathione peroxidase--GSH-Px < 0.139 moli/ml and catalase < 2.20 U/ml). The simultaneous low intake of selenium from the central drinking water supplies in the city of Iasi (0.1-1 g/l) has permitted us to consider necessary the diet supplementation both with foods rich in vitamins with an antioxidant role and with specific medication with selenium, as a protective micro-element.

MEDLINE DUPLICATE 9 L33 ANSWER 21 OF 35

ACCESSION NUMBER: 97107324 MEDLINE

PubMed ID: 8950072 DOCUMENT NUMBER: 97107324

TITLE: Antioxidant status of hypercholesterolemic patients treated

with LDL apheresis.

Lepage S; Bonnefont-Rousselot D; Bruckert E; Bourely B; AUTHOR:

Jaudon M C; Delattre J; Assogba U

Laboratoire de Biochimie, Hopital Pitie-Salpetriere, Paris, CORPORATE SOURCE:

France.

CARDIOVASCULAR DRUGS AND THERAPY, (1996 Nov) 10 (5) 567-71. SOURCE:

Journal code: 8712220. ISSN: 0920-3206.

PUB. COUNTRY: United States

(CLINICAL TRIAL) DOCUMENT TYPE:

(CONTROLLED CLINICAL TRIAL)

Journal; Article; (JOURNAL ARTICLE)

LANGUAGE: English

FILE SEGMENT: Priority Journals

ENTRY MONTH: 199703

ENTRY DATE: Entered STN: 19970313

> Last Updated on STN: 19970313 Entered Medline: 19970303

Oxidation of low density lipoprotein is involved in the pathogenesis of AΒ atherosclerosis. Epidemiological studies suggest a negative correlation between the occurrence of cardiovascular diseases and blood concentrations of lipophilic antioxidants such as vitamins A and E and beta-carotene. Trace elements, such as selenium, zinc, and copper, are involved in the activity of the antioxidant enzymes glutathione peroxidase and superoxide dismutase. The aim of this study was to determine the antioxidant and trace element status of patients with severe hypercholesterolemia who had been treated with dextran-sulphate low-density lipoprotein apheresis in comparison with two control populations, normocholesterolemic subjects and untreated hypercholesterolemic patients. Our results showed that, patients treated with LDL apheresis, compared with normocholesteromic subjects, were not deficient in vitamin E, beta-carotene, and copper, but had lower plasma levels of selenium, zinc, and vitamin A. The low selenium and vitamin A levels were due to the LDL-apheresis treatment, and the hypercholesterolemia might have provoked the low plasma levels of zinc. The study pointed out the potential benefits of supplemental selenium, zinc, and vitamin A in patients being treated with LDL apheresis.

. cardiovascular diseases and blood concentrations of lipophilic ABantioxidants such as vitamins A and E and beta-carotene. Trace elements, such as selenium, zinc, and copper, are involved in the activity of the antioxidant enzymes glutathione peroxidase and superoxide dismutase. The aim of this study was to determine the antioxidant and trace element status of patients with severe hypercholesterolemia who had been treated with dextran-sulphate low-density lipoprotein apheresis in comparison with two control populations, normocholesterolemic subjects and untreated hypercholesterolemic. . . apheresis, compared with normocholesteromic subjects, were not deficient in vitamin E, beta-carotene, and copper, but had lower plasma levels of selenium , zinc, and vitamin A. The low selenium and vitamin A levels were due to the LDL-apheresis treatment, and the hypercholesterolemia might have provoked the low plasma levels of

zinc. The study pointed out the potential benefits of supplemental

selenium, zinc, and vitamin A in patients being treated with LDL apheresis.

L33 ANSWER 22 OF 35 CAPLUS COPYRIGHT 2003 ACS ACCESSION NUMBER: 1995:888305 CAPLUS

DOCUMENT NUMBER: 123:311475

TITLE: Preferential depletion of selenoprotein P in

hypercholesterolemic patients treated by LDL-apheresis AUTHOR (S): Persson-Moschos, M.; Bonnefont-Rousselot, D.; Assoqba,

U.; Bruckert, E.; Jaudon, M. C.; Delattre, J.;

Akesson, B.

Department of Applied Nutrition and Food Chemistry, CORPORATE SOURCE:

University of Lund, P.O. Box 124, Lund, Swed. SOURCE: Clinica Chimica Acta (1995), 240(2), 209-12

CODEN: CCATAR; ISSN: 0009-8981

PUBLISHER: Elsevier DOCUMENT TYPE: Journal LANGUAGE: English

Previously that authors have shown that hypercholesterolemic patients treated by LDL-apheresis had a mean selenium concn. of 0.88 .mu.mol/L, which was lower than control normocholesterolemics. Here, the results indicate that, as a contributory factor for lowered plasma selenium, selenoprotein P is selectively depleted by LDL-apheresis. The affinity of selenoprotein P for sulfated polysaccharides such as dextran sulfate cellulose used in apheresis, is suggested to play a role in the depletion.

Glycoproteins, specific or class

RL: BOC (Biological occurrence); BSU (Biological study, unclassified);

BIOL (Biological study); OCCU (Occurrence)

(selenium-contg., P, selective plasma depletion in humans undergoing LDL-apheresis for hypercholesterolemia)

L33 ANSWER 23 OF 35 BIOSIS COPYRIGHT 2003 BIOLOGICAL ABSTRACTS INC.

ACCESSION NUMBER: 1995:306063 BIOSIS DOCUMENT NUMBER: PREV199598320363

TITLE: Ten-year retrospective on the antioxidant hypothesis of arteriosclerosis: Threshold plasma levels of antioxidant

micronutrients related to minimum cardiovascular risk.

Gey, K. Fred AUTHOR(S):

Inst. Biochem. Mol. Biol., Univ. Berne, Buhlstrasse 28, CORPORATE SOURCE:

CH-3012 Berne Switzerland

SOURCE: Journal of Nutritional Biochemistry, (1995) Vol. 6, No. 4,

pp. 206-236. ISSN: 0955-2863.

DOCUMENT TYPE: General Review

LANGUAGE: English

The antioxidant hypothesis postulates that suboptimal levels of principal antioxidant micronutrients are hitherto underrated risk factors for cardiovascular diseases. Complementary observational data consistently suggest optimal, i.e., potentially protective plasma levels of approximately gt 50 mu-mol/L of vitamin C, gt 30 mu-mol/L of lipid-standardized vitamin E alpha-tocopherol/cholesterol ratio gt 5.2 mu-mol/mmol), and gt 0.4 mu-mol/L beta (gt 0.5 mu-mol/L total)-carotene. Relative risks are doubled at gt 25 to 50% lower values. Suboptimal levels of each factor increase the risk singly, or in combination risk increases multiplicatively. They can be stronger predictors of coronary heart disease than classical risk factors such as hypercholesterolemia and hypertension, at least in Northern Europe. In male Americans, the relative risk of cardiovascular diseases was substantially reduced by daily intake of gt 130 mg of vitamin C, gt 100 IU of vitamin E (100 mg of d,1- or 74 mg of d-alpha-acetyl-tocopherol) in all subjects, and by gt 9 mg of beta-carotene, but only in smokers-in comparison with a suboptimal intake that very probably permits only suboptimal plasma levels. Antioxidant deficits can be avoided by "prudent diets" rich in fruits/vegetables, and net vitamin E (high vitamin E/polyunsaturated fatty acids ratio) as is common in European communities where premature cardiovascular death is low. These essential antioxidants may be crucial components of such protective diets but other, presumably synergistic constituents await evaluation, e.g., carotenoids other than beta-carotene, phenols/bioflavonoids, minerals such as potassium and selenium, fibers, mono- and n-3 polyenic fatty acids, and oxygen-sensitive B vitamins such as folate.

AB. . . in combination risk increases multiplicatively. They can be stronger predictors of coronary heart disease than classical risk factors such as

hypercholesterolemia and hypertension, at least in Northern Europe. In male Americans, the relative risk of cardiovascular diseases was substantially reduced by. . . protective diets but other, presumably synergistic constituents await evaluation, e.g., carotenoids other than beta-carotene, phenols/bioflavonoids, minerals such as potassium and selenium, fibers, mono- and n-3 polyenic fatty acids, and oxygen-sensitive B vitamins such as folate.

=> d ibib abs kwic 15-19

L33 ANSWER 15 OF 35 MEDLINE DUPLICATE 6

ACCESSION NUMBER: 1998442167 MEDLINE

DOCUMENT NUMBER: 98442167 PubMed ID: 9770031

TITLE: [The plasma antioxidant status and trace elements in

patients with familial hypercholesterolemia treated with

LDL-apheresis].

Statut plasmatique en antioxydants et en oligoelements de patients atteints d'hypercholesterolemie familiale traites

par LDL-apherese.

AUTHOR: Delattre J; Lepage S; Jaudon M C; Bruckert E; Assogba U;

Bonnefont-Rousselot D

CORPORATE SOURCE: Laboratoire de Biochimie Metabolique et Clinique, Faculte

de Pharmacie, Hopital Pitie-Salpetriere, Paris.

SOURCE: ANNALES PHARMACEUTIQUES FRANCAISES, (1998) 56 (1) 18-25.

Journal code: 2985176R. ISSN: 0003-4509.

PUB. COUNTRY: France

DOCUMENT TYPE: (CLINICAL TRIAL)

Journal; Article; (JOURNAL ARTICLE)

LANGUAGE: French

FILE SEGMENT: Priority Journals

ENTRY MONTH: 199811

ENTRY DATE: Entered STN: 19990106

Last Updated on STN: 19990106 Entered Medline: 19981110

Oxidation of low density lipoprotein is involved in the pathogenesis of atherosclerosis. Epidemiological studies suggest a negative correlation between the occurrence of cardiovascular diseases and blood concentrations of lipophilic antioxidants such as vitamin A and E and beta-carotene. Trace elements such as selenium, zinc and copper are involved in the activity of antioxidant enzymes: glutathione peroxidase and superoxide dismutase. The aim of this work was to determine the antioxidant and trace elements status of patients with very severe hypercholesterolemia and who were treated by dextran sulphate low density lipoprotein apheresis, in comparison with two control populations: one constituted by normocholesterolemic subjects and the other by hypercholesterolemic patients before treatment. Our results showed that, as compared with normocholesterolemic subjects, patients treated by LDL-apheresis were not deficient in vitamin E, beta-carotene and copper but had low plasma levels of selenium, zinc and vitamin A. The low selenium and vitamin A levels were due to the treatment by LDL-apheresis by itself, while the hypercholesterolemia of these patients might have provoked the low plasma levels of zinc. This study pointed out the interest of a supplement of selenium, zinc and vitamin A in patients treated by LDL-apheresis.

AR cardiovascular diseases and blood concentrations of lipophilic antioxidants such as vitamin A and E and beta-carotene. Trace elements such as selenium, zinc and copper are involved in the activity of antioxidant enzymes: glutathione peroxidase and superoxide dismutase. The aim of this work was to determine the antioxidant and trace elements status of patients with very severe hypercholesterolemia and who were treated by dextran sulphate low density lipoprotein apheresis, in comparison with two control populations: one constituted by. subjects, patients treated by LDL-apheresis were not deficient in vitamin E, beta-carotene and copper but had low plasma levels of selenium , zinc and vitamin A. The low selenium and vitamin A levels were due to the treatment by LDL-apheresis by itself, while the hypercholesterolemia of these patients might have provoked the low plasma levels of zinc. This study pointed out the interest of a supplement of selenium, zinc and vitamin A in patients treated by LDL-apheresis.

ACCESSION NUMBER: 1998:150183 CAPLUS

DOCUMENT NUMBER: 128:158912

TITLE: Pharmaceutical compositions containing ceramides and

metal salts for the treatment of hypercholesterolemia

INVENTOR(S): Shrivastava, Ravi; Lambropoulos, Patrick

PATENT ASSIGNEE(S): Shrivastava, Ravi, Fr.; Hydroxydase Societe des Eaux

Minerales Naturelles et des Laboratoires du Breuil

SOURCE: Fr. Demande, 11 pp.

CODEN: FRXXBL

DOCUMENT TYPE: LANGUAGE: Patent French

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
FR 2747307	A1	19971017	FR 1996-4762	19960411
FR 2747307	B1	19980710		

PRIORITY APPLN. INFO.: FR 1996-4762 19960411

AB Pharmaceutical compns. contg. ceramides and metal salts or vitamin E are useful for the treatment of hypercholesterolemia. Combination of wheat ceramides (4 mg/kg/day) and magnesium oxide (35 mg/kg/day) reduced cholesterol level in hypercholesterolemic rabbits by 47.9% after 4 wk of treatment. Pharmaceutical tablets contained wheat ceramides 100, magnesium oxide 200, and excipients 500 mg/tablet.

IT 59-02-9, .alpha.-Tocopherol 1309-48-4, Magnesium oxide, biological studies 1406-18-4, Vitamin E 7439-89-6D, Iron, salts, biological studies 7439-93-2D, Lithium, salts, biological studies 7439-95-4D, Magnesium, salts, biological studies 7439-96-5D, Manganese, salts, biological studies 7440-02-0D, Nickel, salts, biological studies 7440-21-3D, Silicon, salts, biological studies 7440-48-4D, Cobalt, salts, biological studies 7440-66-6D, Zinc, salts, biological studies 7782-49-2D, Selenium, salts, biological studies

RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(pharmaceutical compns. contg. ceramides and metal salts for treatment of hypercholesterolemia)

L33 ANSWER 17 OF 35 MEDLINE DUPLICATE 7

ACCESSION NUMBER: 1998059634 MEDLINE

DOCUMENT NUMBER: 98059634 PubMed ID: 9397245

TITLE: Effect of doxazosin on endothelial dysfunction in

hypercholesterolemic/antioxidant-deficient rats.

AUTHOR: Raij L; Hayakawa H; Coffee K; Guerra J

CORPORATE SOURCE: Renal Section 111J, Veterans Affairs Medical Center,

Minneapolis, MN 55417, USA.

SOURCE: AMERICAN JOURNAL OF HYPERTENSION, (1997 Nov) 10 (11)

1257-62.

Journal code: 8803676. ISSN: 0895-7061.

PUB. COUNTRY: United States

DOCUMENT TYPE: Journal; Article; (JOURNAL ARTICLE)

LANGUAGE: English

FILE SEGMENT: Priority Journals

ENTRY MONTH: 199801

ENTRY DATE: Entered STN: 19980129

Last Updated on STN: 19980129 Entered Medline: 19980109

AB Hypertension, hypercholesterolemia, atherosclerosis, and coronary heart disease are associated with abnormal endothelium-dependent, nitric oxide-mediated vasorelaxation. In rats, hypercholesterolemia in combination with deficiencies of vitamin E and selenium results in increased endogenous lipid oxidation and endothelial dysfunction. Two hydroxymetabolites of doxazosin, an alpha 1-adrenergic blocking antihypertensive agent, inhibit human lipid oxidation in vitro in a dose-dependent fashion. The present studies were performed to determine the effect of in vivo treatment with doxazosin on endothelial dysfunction in hypercholesterolemic/ antioxidant-deficient rats. Dahl rats were fed 1) a standard diet, 2) a high cholesterol (4%) diet, or 3) a high cholesterol, vitamin E- and selenium -deficient diet. A subgroup of animals in each group were administered doxazosin (3.5 mg/100 g/day) for 16 weeks. In the aortas, vascular

relaxations induced by acetylcholine were significantly decreased (P < .05) in high cholesterol/antioxidant-deficient rats compared with normal and high cholesterol animals. Doxazosin treatment prevented the impairment in endothelium-dependent vascular relaxation in the high cholesterol/antioxidant-deficient group. Vasorelaxation in response to the exogenous nitric oxide donor diethylamine nanoate, which was significantly impaired (P < .05) in aortas from high cholesterol/antioxidant-deficient animals compared with normal and high cholesterol animals, was normalized in aortas from high cholesterol/ antioxidant-deficient animals that had received doxazosin. The antioxidant effect of doxazosin may have therapeutic implications in diseases associated with endothelial dysfunction linked to products of lipid oxidation.

Hypertension, hypercholesterolemia, atherosclerosis, and coronary heart disease are associated with abnormal endothelium-dependent, nitric oxide-mediated vasorelaxation. In rats, hypercholesterolemia in combination with deficiencies of vitamin E and selenium results in increased endogenous lipid oxidation and endothelial dysfunction. Two hydroxymetabolites of doxazosin, an alpha 1-adrenergic blocking antihypertensive agent, inhibit. . . were fed 1) a standard diet, 2) a high cholesterol (4%) diet, or 3) a high cholesterol, vitamin E- and selenium-deficient diet. A subgroup of animals in each group were administered doxazosin (3.5 mg/100 g/day) for 16 weeks. In the aortas, . .

L33 ANSWER 18 OF 35 MEDLINE DUPLICATE 8

ACCESSION NUMBER: 1998249234 MEDLINE

DOCUMENT NUMBER: 98249234 PubMed ID: 9587653

TITLE: Effect of diet induced hypercholesterolemia and

selenium supplementation on nitric oxide synthase

activity.

AUTHOR: Kang B P; Mehta U; Bansal M P

CORPORATE SOURCE: Department of Biophysics, Panjab University, Chandigarh,

India.

SOURCE: ARCHIVES OF PHYSIOLOGY AND BIOCHEMISTRY, (1997 Oct) 105 (6)

603-6.

Journal code: 9510153. ISSN: 1381-3455.

PUB. COUNTRY: Netherlands

DOCUMENT TYPE: Journal; Article; (JOURNAL ARTICLE)

LANGUAGE: English

FILE SEGMENT: Priority Journals

ENTRY MONTH: 199807

ENTRY DATE: Entered STN: 19980811

Last Updated on STN: 19980811 Entered Medline: 19980727

The aim of the present study was to examine the activity of nitric oxide synthase (NOS, EC 1.14.23) in plasma of high fat diet (HFD, 2% cholesterol and 100 g table butter/kg diet) and HFD + selenium (Se, 1 ppm as sodium selenite) fed rabbits for three months. Significant increase in the serum cholesterol and triglyceride levels in HFD fed group was observed. The activity of NOS also increased very significantly. However in Se supplemented animals, there was a significant reduction in serum cholesterol as well as in plasma NOS activity relative to HFD fed animals. It is concluded that the protective effect of Se on HFD induced NOS activity acts probably through its antioxidant/inhibitory action.

TI Effect of diet induced hypercholesterolemia and selenium supplementation on nitric oxide synthase activity.

L33 ANSWER 19 OF 35 CAPLUS COPYRIGHT 2003 ACS ACCESSION NUMBER: 1997:592217 CAPLUS

DOCUMENT NUMBER: 127:277540

TITLE: Effect of selenium deficiency on hepatic lipid and

lipoprotein metabolism in the cat

AUTHOR(S): Nassir, F.; Moundras, C.; Bayle, D.; Serougne, C.;

Gueux, E.; Rock, E.; Rayssiguier, Y.; Mazur, A. Centre REcherche Nutrition Humaine, Unite Maladies

Metaboliques Micronutriments, Saint-Genes-Champanelle,

63122, Fr.

SOURCE: British Journal of Nutrition (1997), 78(3), 493-500

CODEN: BJNUAV; ISSN: 0007-1145

PUBLISHER: CAB International

DOCUMENT TYPE: Journal LANGUAGE: English

CORPORATE SOURCE:

AB Since exptl. Se deficiency results in a significant increase in plasma

cholesterol concn. the present investigation was undertaken to assess further the influence of this deficiency on the expression of proteins involved in hepatic lipid metab. Se deficiency was induced by feeding weanling male Wistar rats on a deficient diet for 6 wk. Hypercholesterolemia assocd. with Se deficiency was related to increased 3-hydroxy-3-methylglutaryl-coA (HMG-CoA) reductase (EC 1.1.1.34) activity in liver microsomes as compared with control animals. Hepatic lipoprotein receptor levels (LDL-receptor and HDL-binding proteins, HB1 and HB2) were not significantly affected by Se deficiency, as assessed by immunoblotting. Plasma triacylglycerol concns. tended to decrease in Se-deficient rats in concert with their reduced post-Triton secretion. There was no significant effect of Se deficiency on the hepatic synthesis of apolipoproteins. These results point to the need for further investigations into the mechanism related to the increased activity of HMG-CoA reductase and the enhanced cholesterogenesis in the liver of Se-deficient rats likely to result from this.

TΥ Cat (Felis catus)

Hypercholesterolemia

Liver

AUTHOR (S):

PUBLISHER:

(effect of selenium deficiency on hepatic lipid and lipoprotein metab. in the cat)

=> d ibib abs kwic 10-14

L33 ANSWER 10 OF 35 CAPLUS COPYRIGHT 2003 ACS ACCESSION NUMBER: 1999:386550 CAPLUS

DOCUMENT NUMBER: 131:183269

TITLE: Relationship between hypercholesterolemia, endothelial

dysfunction and hypertension

Hayakawa, Hiroshi; Raij, Leopoldo Department of Medicine, Veterans Affairs Medical CORPORATE SOURCE:

Center and University of Minnesota Medical School,

Minneapolis, MN, 55417, USA

Journal of Hypertension (1999), 17(5), 611-619 SOURCE:

CODEN: JOHYD3; ISSN: 0263-6352 Lippincott Williams & Wilkins

DOCUMENT TYPE: Journal LANGUAGE: English

We have previously shown that in the rat a diet high in cholesterol and deficient in vitamin E and selenium results in hypercholesterolemia and increased lipid oxidn. We utilized this model to det. whether rats given this diet develop impaired endothelium-dependent relaxation mediated by nitric oxide (NO) in mesenteric and in renal vessels. In addn., we tested whether the impairment is due to (i) decreased endothelial NO synthase activity, (ii) increased NO inactivation and/or (iii) increased prodn. of the endothelium-derived constricting factors thromboxane A2/prostaglandin H2 and endothelin-1. We also investigated whether endothelial dysfunction induced by dyslipidemia increases the sensitivity for the development of hypertension in response to high dietary salt. Male Dahl salt-sensitive (DSS) rats were divided into three groups and received a std. diet (control group), a high (4%) cholesterol diet (HChol), or a high cholesterol diet deficient in the anti-oxidants vitamin E and selenium (HChol-Def). The NaCl content of these diets was 0.5%. After 18 wk we studied endothelium-dependent relaxation in response to acetylcholine (ACh) in aortas and in isolated perfused prepns. of mesenteric arteries and kidneys. In some expts., ifetroban, a thromboxane A2/prostaglandin H2 receptor antagonist, was added to the organ bath or the perfusion buffer. Vascular responses to endothelin-1 as well as to BQ-123, an endothelin A receptor blocker, were studied in the isolated perfused kidneys. In addn., two extra groups of rats were fed a diet high in sodium chloride (2%): one of the groups received the normal cholesterol diet whereas the other group received the diet high in cholesterol and deficient in vitamin E and selenium. Compared to normocholesterolemic rats, responses to ACh were significantly impaired in aortas, mesenteric arteries and kidneys of HChol-Def rats (P<0.01). Endothelial NO synthase activity (conversion of [14C] L-arginine to [14C] L-citrulline) was similar in aortas of control, HChol and HChol-Def rats; thus suggesting that impaired endothelium-dependent relaxation in the HChol-Def rats was not due to decreased cNOS catalytic activity. Ifetroban improved the impaired endothelium-dependent relaxation in mesenteric vessels, but not in aortas and kidneys. Endothelin-1 (ET-1:10-13-10-11 mol/1) elicited NO-mediated

relaxations in kidneys of control rats but not in kidneys of HChol-Def; blockade of ET-1 with BQ-123, an ETA receptor blocker, did not improve NO-mediated relaxation of HChol-Def. Despite impaired endothelium-dependent relaxation in renal and mesenteric vessels, HChol-Def DSS rats failed to develop hypertension (systolic blood pressure 144.+-.1 in control and 150.+-.2 mmHg in HChol-Def) but manifested a significant increase in sensitivity to the pressor effects of a high (2% NaCl) dietary salt content during the initial 10 wk of the study, although the final blood pressure at 18 wk was similar in both groups. These studies support the notion that (i) products of lipid oxidn. may reduce NO bioactivity without affecting endothelial NO synthase mass or catalytic activity, (ii) the mechanisms involved in the endothelial dysfunction induced by hypercholesterolemia and oxidized lipids may differ among vascular beds, and (iii) decreased NO bioavailability does not necessarily result in systemic hypertension, but it may enhance the sensitivity to the hypertensinogenic effect of dietary salt.

THERE ARE 42 CITED REFERENCES AVAILABLE FOR THIS REFERENCE COUNT: 42 RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

We have previously shown that in the rat a diet high in cholesterol and deficient in vitamin E and selenium results in hypercholesterolemia and increased lipid oxidn. We utilized this model to det. whether rats given this diet develop impaired endothelium-dependent relaxation mediated by nitric oxide (NO) in mesenteric and in renal vessels. In addn., we tested whether the impairment is due to (i) decreased endothelial NO synthase activity, (ii) increased NO inactivation and/or (iii) increased prodn. of the endothelium-derived constricting factors thromboxane A2/prostaglandin H2 and endothelin-1. We also investigated whether endothelial dysfunction induced by dyslipidemia increases the sensitivity for the development of hypertension in response to high dietary salt. Male Dahl salt-sensitive (DSS) rats were divided into three groups and received a std. diet (control group), a high (4%) cholesterol diet (HChol), or a high cholesterol diet deficient in the anti-oxidants vitamin E and selenium (HChol-Def). The NaCl content of these diets was 0.5%. After 18 wk we studied endothelium-dependent relaxation in response to acetylcholine (ACh) in aortas and in isolated perfused prepns. of mesenteric arteries and kidneys. In some expts., ifetroban, a thromboxane A2/prostaglandin H2 receptor antagonist, was added to the organ bath or the perfusion buffer. Vascular responses to endothelin-1 as well as to BQ-123, an endothelin A receptor blocker, were studied in the isolated perfused kidneys. In addn., two extra groups of rats were fed a diet high in sodium chloride (2%): one of the groups received the normal cholesterol diet whereas the other group received the diet high in cholesterol and deficient in vitamin E and selenium. Compared to normocholesterolemic rats, responses to ACh were significantly impaired in aortas, mesenteric arteries and kidneys of HChol-Def rats (P<0.01). Endothelial NO synthase activity (conversion of [14C] L-arginine to [14C] L-citrulline) was similar in aortas of control, HChol and HChol-Def rats; thus suggesting that impaired endothelium-dependent relaxation in the HChol-Def rats was not due to decreased cNOS catalytic activity. Ifetroban improved the impaired endothelium-dependent relaxation in mesenteric vessels, but not in aortas and kidneys. Endothelin-1 (ET-1:10-13-10-11 mol/1) elicited NO-mediated relaxations in kidneys of control rats but not in kidneys of HChol-Def; blockade of ET-1 with BQ-123, an ETA receptor blocker, did not improve NO-mediated relaxation of HChol-Def. Despite impaired endothelium-dependent relaxation in renal and mesenteric vessels, HChol-Def DSS rats failed to develop hypertension (systolic blood pressure 144.+-.1 in control and 150.+-.2 mmHg in HChol-Def) but manifested a significant increase in sensitivity to the pressor effects of a high (2% NaCl) dietary salt content during the initial 10 wk of the study, although the final blood pressure at 18 wk was similar in both groups. These studies support the notion that (i) products of lipid oxidn. may reduce NO bioactivity without affecting endothelial NO synthase mass or catalytic activity, (ii) the mechanisms involved in the endothelial dysfunction induced by hypercholesterolemia and oxidized lipids may differ among vascular beds, and (iii) decreased NO bioavailability does not necessarily result in systemic hypertension, but it may enhance the sensitivity to the hypertensinogenic effect of dietary salt. IT Diet

(with high cholesterol and deficient in vitamin E and selenium ; relationship between hypercholesterolemia, endothelial dysfunction and hypertension in rats)

IT 1406-18-4, Vitamin E 7782-49-2, Selenium, biological studies RL: BPR (Biological process); BSU (Biological study, unclassified); BIOL (Biological study); PROC (Process)

(deficient, diet; relationship between hypercholesterolemia, endothelial dysfunction and hypertension in rats)

L33 ANSWER 11 OF 35 MEDLINE

ACCESSION NUMBER: 1999459117 MEDLINE

DOCUMENT NUMBER: 99459117 PubMed ID: 10527957

TITLE: Acute effects of LDL-apheresis on cholesterol oxidation

products and antioxidants in plasma and lipoproteins of

patients with familial hypercholesterolemia.

AUTHOR: Linseisen J; Wilhelm M; Hoffmann J; Hailer S; Keller C;

Wolfram G

CORPORATE SOURCE: Institut fur Ernahrungswissenschaft der TU Munchen, D-85350

Freising-Weihenstephan, Germany...

linseisen@weihenstephan.de

SOURCE: EUROPEAN JOURNAL OF MEDICAL RESEARCH, (1999 Oct 15) 4 (10)

433-41.

Journal code: 9517857. ISSN: 0949-2321.

PUB. COUNTRY: GERMANY: Germany, Federal Republic of

DOCUMENT TYPE: Journal; Article; (JOURNAL ARTICLE)

LANGUAGE: English

FILE SEGMENT: Priority Journals

ENTRY MONTH: 199912

ENTRY DATE: Entered STN: 20000113

Last Updated on STN: 20000113 Entered Medline: 19991202

Regular LDL-apheresis treatment of hypercholesterolemic patients has proven to reduce the formation of atherosclerotic lesions. Regarding the underlying mechanisms, cholesterol oxidation products (COP) may play a detrimental role. Therefore, COP levels were determined before and after regular LDL-apheresis treatment in ten patients with familial hypercholesterolemia. - The patients had approximately twofold elevated plasma and LDL COP concentrations on the average as compared to healthy subjects. LDL-apheresis treatment efficiently removed COP from the circulation. As a consequence of a smaller reduction of the COP content (-52 %) than of the total cholesterol content (-71 %) in LDL, the LDL COP: cholesterol ratio increased. Lipid-soluble antioxidants in the plasma of the hypercholesterolemics decreased to a comparable extent as did plasma lipids. In contrast to nearly stable vitamin C concentrations, plasma selenium concentrations also decreased, resulting altogether in a decreased but still normal serum total antioxidant capacity. - In conclusion, LDL-apheresis treatment effectively reduced potentially atherogenic COP from the plasma. With normal plasma antioxidant concentrations before LDL-apheresis in long-term treated hypercholesterolemics, the observed acute decrease in lipid-soluble antioxidants and selenium by treatment seems not to be as meaningful. The higher LDL COP:cholesterol ratio after treatment needs further elucidation.

AB . . . play a detrimental role. Therefore, COP levels were determined before and after regular LDL-apheresis treatment in ten patients with familial hypercholesterolemia. - The patients had approximately twofold elevated plasma and LDL COP concentrations on the average as compared to healthy subjects. . . the hypercholesterolemics decreased to a comparable extent as did plasma lipids. In contrast to nearly stable vitamin C concentrations, plasma selenium concentrations also decreased, resulting altogether in a decreased but still normal serum total antioxidant capacity. - In conclusion, LDL-apheresis treatment. . . plasma. With normal plasma antioxidant concentrations before LDL-apheresis in long-term treated hypercholesterolemics, the observed acute decrease in lipid-soluble antioxidants and selenium by treatment seems not to be as meaningful. The higher LDL COP:cholesterol ratio after treatment needs further elucidation.

L33 ANSWER 12 OF 35 BIOSIS COPYRIGHT 2003 BIOLOGICAL ABSTRACTS INC.

ACCESSION NUMBER: 1998:261078 BIOSIS DOCUMENT NUMBER: PREV199800261078

TITLE: Effect of diet induced hypercholesterolemia and selenium supplementation on nitric oxide synthase

activity.

AUTHOR(S): Kang, B. P. S.; Mehta, U.; Bansal, M. P. (1)

CORPORATE SOURCE: (1) Dep. Biophys., Panjab Univ., Chandigarh 160 014 India SOURCE: Archives of Physiology and Biochemistry, (Oct., 1998) Vol.

105, No. 6, pp. 603-606.

ISSN: 1381-3455.

DOCUMENT TYPE: Article LANGUAGE: English

AB The aim of the present study was to examine the activity of nitric oxide synthase (NOS, EC 1.14.23) in plasma of high fat diet (HFD, 2% cholesterol and 100 g table butter/kg diet) and HFD + selenium (Se, 1 ppm as sodium selenite) fed rabbits for three months. Significant increase in the serum cholesterol and triglyceride levels in HFD fed group was observed. The activity of NOS also increased very significantly. However in Se supplemented animals, there was a significant reduction in serum cholesterol as well as in plasma NOS activity relative to HFD fed animals. It is concluded that the protective effect of Se on HFD induced NOS activity acts probably through its antioxidant/inhibitory action.

Effect of diet induced hypercholesterolemia and selenium ${\tt supplementation} \ {\tt on} \ {\tt nitric} \ {\tt oxide} \ {\tt synthase} \ {\tt activity}.$

L33 ANSWER 13 OF 35 CAPLUS COPYRIGHT 2003 ACS ACCESSION NUMBER: 1998:560601 CAPLUS

DOCUMENT NUMBER: 129:259722

TITLE: Vitamin E combined with selenium inhibits atherosclerosis in hypercholesterolemic rabbits independently of effects on plasma cholesterol

concentrations

Schwenke, Dawn C.; Behr, Stephen R. AUTHOR (S):

CORPORATE SOURCE: Department of Pathology, Wake Forest University School

of Medicine, Winston-Salem, NC, 27157-1072, USA

SOURCE: Circulation Research (1998), 83(4), 366-377

CODEN: CIRUAL; ISSN: 0009-7330

PUBLISHER: Williams & Wilkins

DOCUMENT TYPE: Journal LANGUAGE: English

Combining vitamin E with vitamin C and/or selenium could inhibit atherosclerosis more effectively than vitamin E alone. Rabbits were fed a control atherogenic diet or the atherogenic diet supplemented with vitamin E, vitamins E + C, vitamin E + Se, vitamins E + C + Se, or probucol (pos. control). The daily supplements were 146 IU vitamin E, 791 mg vitamin C, 22 .mu.g Se, or 406 mg probucol. Vitamin C did not influence the atherosclerosis process. After 22 wk of treatment, rank order of aortic atherosclerosis was control > vitamin E (with or without vitamin C) > vitamin E + selenium (with or without vitamin C) > probucol. The antioxidant treatment decreased the aortic cholesterol concns. 21-56, 29-86, and 19-75% for the aortic arch, descending thoracic aorta, and abdominal aorta, resp., with slightly greater decreases in areas of atherosclerotic lesions. Some treatments reduced blood plasma cholesterol concns., but none altered the distribution of cholesterol in lipoproteins. When cor. for differences in blood plasma cholesterol concns., the aortic cholesterol concns. were reduced up to 72% by the antioxidant treatments, with equal redns. by vitamin E + Se and by probucol. The aortic .alpha.-tocopherol standardized by aortic cholesterol as a measure of aortic lipids was lower in the abdominal aorta than in the aortic arch of rabbits not given .alpha.-tocopherol, and increased relatively more in the abdominal aorta than in the aortic arch with .alpha.-tocopherol supplementation. Thus, vitamin E + Se inhibited atherosclerosis as effectively as equally hypocholesterolemic doses of probucol by a mechanism(s) independent of the effects on blood plasma and lipoprotein cholesterol concns. The tendency for greater efficacy of antioxidant treatments in the abdominal aorta than aortic arch may be related to the lower concns. of .alpha.-tocopherol in the abdominal aorta of nonsupplemented rabbits.

REFERENCE COUNT: THERE ARE 77 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

Antioxidants

Atherosclerosis Blood plasma

Hypercholesterolemia

Nutrition, animal

(dietary vitamins E plus C and selenium inhibit atherosclerosis in hypercholesterolemic rabbits independently of effects on blood cholesterolemia)

L33 ANSWER 14 OF 35 MEDLINE

ACCESSION NUMBER: 1998340157 MEDLINE DUPLICATE 5

DOCUMENT NUMBER:

98340157 PubMed ID: 9675557

TITLE:

Selenium supplementation and diet induced

hypercholesterolemia in the rat: changes in lipid

levels, malonyldialdehyde production and the nitric oxide

synthase activity.

Kanq B P; Bansal M P; Mehta U AUTHOR:

CORPORATE SOURCE:

Department of Biophysics, Panjab University Chandigarh,

India.

SOURCE:

GENERAL PHYSIOLOGY AND BIOPHYSICS, (1998 Mar) 17 (1) 71-8.

Journal code: 8400604. ISSN: 0231-5882.

PUB. COUNTRY: DOCUMENT TYPE: Slovakia

Journal; Article; (JOURNAL ARTICLE)

LANGUAGE: FILE SEGMENT: English

Priority Journals

ENTRY MONTH:

ENTRY DATE:

199810 Entered STN: 19981020

Last Updated on STN: 19981020

Entered Medline: 19981005

Male Sprague Dawley rats were divided into three groups, viz (I) Controls, (II) High fat diet (HFD) fed, (III) HFD fed+selenium supplemented. After three months of treatment, there were significant increases in serum cholesterol and triglycerides in HFD fed group as compared to control. However, in Se supplemented group, the levels of serum cholesterol and triglycerides were significantly less as compared to group II. Selenium-dependent glutathione peroxidase (GSH-Px) activity in the liver and the aorta increased significantly in HFD fed animals and also showed additional significant increase on selenium supplementation. Malonyldialdehyde (MDA) concentrations in serum, liver and aorta and the activity of nitric oxide synthase (NOS; evident from reactive nitrogen intermediates and citrulline levels) in plasma showed significant increases in HFD fed group. However, supplementation of selenium led to a significant reduction in the levels of these parameters vis-a-vis HFD fed animals except in MDA levels in the serum and the liver where this decrease was non-significant. The important finding of this study is that selenium supplementation modulates the sequences favoring pathogenesis of atherosclerosis.

Selenium supplementation and diet induced hypercholesterolemia in the rat: changes in lipid levels, malonyldialdehyde production and the nitric oxide synthase activity.

=> s phendimetrazinum

L34 2 PHENDIMETRAZINUM

=> fil reg

COST IN U.S. DOLLARS SINCE FILE TOTAL ENTRY SESSION FULL ESTIMATED COST 37.92 190.02 DISCOUNT AMOUNTS (FOR QUALIFYING ACCOUNTS) SINCE FILE TOTAL ENTRY SESSION CA SUBSCRIBER PRICE -3.91 -6.51

FILE 'REGISTRY' ENTERED AT 11:29:24 ON 08 JAN 2003 USE IS SUBJECT TO THE TERMS OF YOUR STN CUSTOMER AGREEMENT. PLEASE SEE "HELP USAGETERMS" FOR DETAILS. COPYRIGHT (C) 2003 American Chemical Society (ACS)

Property values tagged with IC are from the ZIC/VINITI data file provided by InfoChem.

STRUCTURE FILE UPDATES: 7 JAN 2003 HIGHEST RN 478336-86-6 DICTIONARY FILE UPDATES: 7 JAN 2003 HIGHEST RN 478336-86-6

TSCA INFORMATION NOW CURRENT THROUGH MAY 20, 2002

Please note that search-term pricing does apply when conducting SmartSELECT searches.

Crossover limits have been increased. See HELP CROSSOVER for details.

Experimental and calculated property data are now available. See HELP PROPERTIES for more information. See STNote 27, Searching Properties

in the CAS Registry File, for complete details: http://www.cas.org/ONLINE/STN/STNOTES/stnotes27.pdf

=> s phendimetrazinum

O PHENDIMETRAZINUM

L35 0 PHENDIMETRAZINUM

=> s phendimetrazine

L36 6 PHENDIMETRAZINE

=> s phendimetrazine/cn

L37 1 PHENDIMETRAZINE/CN

=> s phendimetrazine/cn or phendimetrazine/crn

1 PHENDIMETRAZINE/CN

0 PHENDIMETRAZINE/CRN

L38 1 PHENDIMETRAZINE/CN OR PHENDIMETRAZINE/CRN

=> FIL MEDL CAPL BIOSIS USPATF

COST IN U.S. DOLLARS
SINCE FILE TOTAL
ENTRY SESSION
FULL ESTIMATED COST 17.68 207.70

DISCOUNT AMOUNTS (FOR QUALIFYING ACCOUNTS)

SINCE FILE TOTAL
ENTRY SESSION

CA SUBSCRIBER PRICE 0.00 -6.51

FILE 'MEDLINE' ENTERED AT 11:30:36 ON 08 JAN 2003

FILE 'CAPLUS' ENTERED AT 11:30:36 ON 08 JAN 2003 USE IS SUBJECT TO THE TERMS OF YOUR STN CUSTOMER AGREEMENT. PLEASE SEE "HELP USAGETERMS" FOR DETAILS. COPYRIGHT (C) 2003 AMERICAN CHEMICAL SOCIETY (ACS)

FILE 'BIOSIS' ENTERED AT 11:30:36 ON 08 JAN 2003 COPYRIGHT (C) 2003 BIOLOGICAL ABSTRACTS INC.(R)

FILE 'USPATFULL' ENTERED AT 11:30:36 ON 08 JAN 2003 CA INDEXING COPYRIGHT (C) 2003 AMERICAN CHEMICAL SOCIETY (ACS)

=> s 138 or phendimetrazine

L39 463 L38 OR PHENDIMETRAZINE

=> s 111 and 139

L40 30 L11 AND L39

=> s 111 (S) 139

L41 0 L11 (S) L39

=> dup rem 140

PROCESSING COMPLETED FOR L40

L42 30 DUP REM L40 (0 DUPLICATES REMOVED)

=> d ibib abs kwic 27-30

L42 ANSWER 27 OF 30 USPATFULL

ACCESSION NUMBER: 1998:98932 USPATFULL

TITLE: DHA-pharmaceutical agent conjugates of taxanes
INVENTOR(S): Shashoua, Victor E., Brookline, MA, United States
Swindell, Charles S., Merion, PA, United States
Webb, Nigel L., Bryn Mawr, PA, United States

Bradley, Matthews O., Laytonsville, MD, United States

PATENT ASSIGNEE(S): Neuromedica, Inc., Conshohocken, PA, United States

(U.S. corporation)

DOCUMENT TYPE: Utility FILE SEGMENT: Granted

PRIMARY EXAMINER: Jarvis, William R. A.

LEGAL REPRESENTATIVE: Wolf, Greenfield & Sacks, P.C.

NUMBER OF CLAIMS: 12 EXEMPLARY CLAIM: NUMBER OF DRAWINGS: 27 Drawing Figure(s); 14 Drawing Page(s) 2451 LINE COUNT: CAS INDEXING IS AVAILABLE FOR THIS PATENT. The invention provides conjugates of cis-docosahexaenoic acid and taxanes useful in treating cell proliferative disorders. Conjugates of paclitaxel and docetaxel are preferred. CAS INDEXING IS AVAILABLE FOR THIS PATENT. . . . They further are useful in treating diabetes and its complications, excess acid secretion, cardiovascular conditions involving cholesterol (e.g., hyperlipidemia and hypercholesterolemia), diarrhea, ovarian diseases (e.g. endometriosis, ovarian cysts, etc.) and as contraceptive agents. Other conditions treatable according to the invention will. . . DETD Appetite suppressant: Dexfenfluramine Hydrochloride; Phendimetrazine Tartrate; Phentermine Hydrochloride. . . . They further are useful in treating diabetes and its DETD complications, excess acid secretion, cardiovascular conditions involving cholesterol (e.g., hyperlipidemia and hypercholesterolemia), diarrhea, ovarian diseases (e.g. endometriosis, ovarian cysts, etc.) and as contraceptive agents. L42 ANSWER 28 OF 30 USPATFULL ACCESSION NUMBER: 1998:98904 USPATFULL TITLE: Method and composition for treating obesity and related disorders in animals comprising dehydroepiandrosterone (DHEA), or a derivative thereof, and an anorectic agent INVENTOR (S): Svec, Frank, Metairie, LA, United States Porter, Johnny, Metairie, LA, United States Louisiana State University Medical Center Foundation, PATENT ASSIGNEE(S): New Orleans, LA, United States (U.S. corporation) NUMBER KIND DATE US 5795880 PATENT INFORMATION: 19980818 APPLICATION INFO.: US 1996-774521 19961230 (8) DOCUMENT TYPE: Utility FILE SEGMENT: Granted PRIMARY EXAMINER: Weddington, Kevin E. LEGAL REPRESENTATIVE: Finnegan, Henderson, Farabow, Garrett & Dunner, L.L.P. NUMBER OF CLAIMS: 37 EXEMPLARY CLAIM: 27 Drawing Figure(s); 13 Drawing Page(s) NUMBER OF DRAWINGS: LINE COUNT: 888 CAS INDEXING IS AVAILABLE FOR THIS PATENT. The invention describes a method and composition for treating obesity or related disorders in animals using an anorectic agent and dehydroepiandrosterone (DHEA). The composition effectively diminishes caloric intake, may alter metabolism, weight gain, or a combination CAS INDEXING IS AVAILABLE FOR THIS PATENT. SUMM . . . following anorectic agents for the treatment of obesity and related disorders in animals: phenylpropanolamine hydrochloride (HCL), fenfluramine HCl, phentermine HCl, phendimetrazine tartrate, mazindol, diethylpropion HCl, fluoxetine HCl, and sibutramine HCl. The anorectic drugs used in this invention include phenylpropanolamine SUMM HCL, fenfluramine HCl, phentermine HCl, phendimetrazine tartrate, mazindol, diethylpropion HCl, fluoxetine HCl, and sibutramine hydrochloride. Tachyphylaxis and tolerance have been demonstrated with all drugs of this. SUMM Phendimetrazine tartrate, chemically known as (+)-3,4-dimethyl-2-phenylmorpholine tartrate, is another anorectic drug having an effect on appetite and the central nervous system.. . . DETD The anorectic agents that can be employed in the invention include phenylpropanolamine HCL, fenfluramine HCl, phentermine HCl, phendimetrazine tartrate, mazindol, diethylpropion HCl, fluoxetine HCl, and sibutramine hydrochloride. . . . 255 (Endocrinol Metab 18):E229-E235 (1988); and Alarrayed et DETD al., "Is There a Role for the Adrenals in the Development of

Hypercholesterolemia in Zucker Fatty Rats, " Am J. Physiol, 263

(Endocrinol Metab 26):E287-E295 (1992). The animals have two phenotypes, obese and lean.. .

DETD . . . a 28 day period. Given the known activities of DHEA, and the anorectic agents phenylpropanolamine HCL, fenfluramine HCl, phentermine HCl, phendimetrazine tartrate, mazindol, diethylpropion HCl, fluoxetine HCl, and sibutramine hydrochloride administered individually, the effect of the combination of the drugs is. . .

CLM What is claimed is:

. at least one anorectic agent is selected from the group consisting of phenylpropanolamine hydrochloride (HCl), fenfluramine hydrochloride (HCl), phentermine HCl, phendimetrazine tartrate, mazindol, diethylpropion HCl, fluoxetine HCl, and sibutramine hydrochloride.

. . at least one anorectic agent is selected from the group consisting of phenylpropanolamine hydrochloride (HCl), fenfluramine hydrochloride (HCl), phentermine HCl, phendimetrazine tartrate, mazindol, diethylpropion HCl, fluoxetine HCl, and sibutramine hydrochloride.

L42 ANSWER 29 OF 30 USPATFULL

ACCESSION NUMBER: 97:7901 USPATFULL

TITLE: Method for treatment or prevention of obesity INVENTOR(S): Clark, Ross G., Pacifica, CA, United States

PATENT ASSIGNEE(S): Genentech, Inc., South San Francisco, CA, United States

(U.S. corporation)

	NUMBER	KIND DATE	
PATENT INFORMATION:	US 5597797	19970128	
	WO 9118621	19911212	
APPLICATION INFO.:	US 1993-150090	19931119	(8)
	WO 1993-US10259	19931026	
		19931119	PCT 371 date
		19931119	PCT 102(e) date

DOCUMENT TYPE: Utility FILE SEGMENT: Granted

PRIMARY EXAMINER: Schain, Howard E.
ASSISTANT EXAMINER: Touzeau, P. Lynn
LEGAL REPRESENTATIVE: Hasak, Janet E.

NUMBER OF CLAIMS: 20 EXEMPLARY CLAIM: 1

NUMBER OF DRAWINGS: 20 Drawing Figure(s); 20 Drawing Page(s)

LINE COUNT: 2197

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

A method is disclosed for treating obese mammals or preventing obesity from occurring in mammals. This method involves administering to the mammal an effective amount of growth hormone in combination with an effective amount of IGF-I. Preferably, the growth hormone is given so as to have a maintained, continual therapeutically effective presence in the blood, such as by continuous infusion or frequent injections, or by use of a long-acting formulation.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

SUMM . . . ovarian disease, dermatological disorders such as infections, varicose veins, Acanthosis nigricans, and eczema, exercise intolerance, diabetes mellitus, insulin resistance, hypertension, hypercholesterolemia, cholelithiasis, osteoarthritis, orthopedic injury, thromboembolic disease, cancer, and coronary heart disease.

Rissanen et al., British Medical Journal, 301: 835-837 (1990).

DETD

DETD . . . of obesity, such as, e.g., arteriosclerosis, Type II diabetes, polycystic ovarian disease, cardiovascular diseases, osteoarthritis, dermatological disorders, hypertension, insulin resistance, hypercholesterolemia, hypertriglyceridemia, and cholelithiasis.

DETD . . . halogenate; cinchocaine; chlorpromazine; appetite-suppressing drugs acting on noradrenergic neurotransmitters such as mazindol and derivatives of phenethylamine, e.g., phenylpropanolamine, diethylpropion, phentermine, phendimetrazine, benzphetamine, amphetamine, methamphetamine, and phenmetrazine; drugs acting on serotonin neurotransmitters such as fenfluramine, tryptophan, 5-hydroxytryptophan, fluoxetine, and sertraline; centrally active. . .

L42 ANSWER 30 OF 30 USPATFULL

ACCESSION NUMBER: 96:53303 USPATFULL

TITLE: Method and composition for treating obesity comprising

dehydroepiandrosterone (DHEA), or a derivative thereof,

and an anorectic agent

INVENTOR(S): Svec, Frank, Metairie, LA, United States

Porter, Johnny, Metairie, LA, United States

PATENT ASSIGNEE(S): Louisiana State Univ. Medical Center Foundation, New

Orleans, LA, United States (U.S. corporation)

NUMBER KIND DATE

PATENT INFORMATION: US 5527788 19960618 APPLICATION INFO.: US 1994-184191 19940118 (8)

DOCUMENT TYPE: Utility FILE SEGMENT: Granted

PRIMARY EXAMINER: Cintins, Marianne M. ASSISTANT EXAMINER: Weddington, Kevin E.

LEGAL REPRESENTATIVE: Finnegan, Henderson, Farabow, Garrett & Dunner

NUMBER OF CLAIMS: 23 EXEMPLARY CLAIM: 1

NUMBER OF DRAWINGS: 27 Drawing Figure(s); 10 Drawing Page(s)

LINE COUNT: 799

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

AB The invention describes a method and composition for treating obesity or related disorders in animals using an anorectic agent and dehydroepiandrosterone (DHEA). The composition effectively diminishes caloric intake, may alter metabolism, weight gain, or a combination thereof.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

SUMM . . . of the following anorectic agents for the treatment of obesity and related disorders in animals: fenfluramine hydrochloride (HCl), phentermine HCl, phendimetrazine tartrate, mazindol, diethylpropion HCl, and fluoxetine HCl. DHEA has been evaluated for its ability to modify food intake and/or weight. . .

SUMM The anorectic drugs used in this invention include fenfluramine HCl, phentermine HCl, phendimetrazine tartrate, mazindol, diethylpropion HCl, and fluoxetine HCl. Tachyphylaxis and tolerance have been demonstrated with all drugs of this class used.

SUMM Phendimetrazine tartrate, chemically known as

(+) -3,4-dimethyl-2-phenylmorpholine tartrate, is another anorectic drug having an effect on appetite and the central nervous system.. . .

DETD The anorectic agents that can be employed in the invention include fenfluramine hydrochloride (HCl), phentermine HCl, phendimetrazine tartrate, mazindol, diethylpropion HCl, and fluoxetine HCl.

DETD . . . 255 (Endocrinol Metab 18):E229-E235 (1988); and Alarrayed et al., "Is There a Role for the Adrenals in the Development of Hypercholesterolemia in Zucker Fatty Rats," Am J. physiol, 263. (Endocrinol Metab 26):E287-E295 (1992). The animals have two phenotypes, obese and lean.. . .

DETD . . . develope over a 28 day period. Given the known activities of DHEA, and the anorectic agents fenfluramine HCl, phentermine HCl, phendimetrazine tartrate, mazindol, diethylpropion HCl, and fluoxetine HCl administered individually, the effect of the combination of the drugs is striking. It. . .

CLM What is claimed is:

. a derivative thereof and at least one anorectic agent selected from the group consisting of fenfluramine hydrochloride (HCl), phentermine HCl, phendimetrazine tartrate, mazindol, diethylpropion HCl, and fluoxetine HCl, and further comprising the administration of a glucocorticoid.

. a derivative thereof and at least one anorectic agent selected from the group consisting of fenfluramine hydrochloride (HCl), phentermine HCl, phendimetrazine tartrate, mazindol, diethylpropion HCl, and fluoxetine HCl. TITLE: Spiro-azacyclic derivatives and their use as

therapeutic agents

INVENTOR(S): Kulagowski, Janusz Jozef, Sawbridgeworth, United

Kingdom

Raubo, Piotr Antoni, Bishops Stortford, United Kingdom Swain, Christopher John, Duxford, United Kingdom Thomson, Christopher George, Sawbridgeworth, United

Kingdom

PATENT ASSIGNEE(S): Merck Sharp & Dohme Ltd., Hertfordshire, United Kingdom

(non-U.S. corporation)

19991118 PCT 371 date 19991118 PCT 102(e) date

NUMBER DATE

PRIORITY INFORMATION: GB 1997-11114 19970529

DOCUMENT TYPE: Utility
FILE SEGMENT: Granted
PRIMARY EXAMINER: Rotman, Alan L.

ASSISTANT EXAMINER: Desai, Rita

LEGAL REPRESENTATIVE: Thies, J. Eric, Rose, David L.

NUMBER OF CLAIMS: 19
EXEMPLARY CLAIM: 1
LINE COUNT: 3494

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

AB Substituted spiro-azacyclic derivatives of structural formula I are tachykinin receptor antagonists of use, for example, in the treatment of pain, inflammation, migraine, emesis and posttherpetic neuralgia

Wherein A is a pyridyl, X is --CH2--, Y is --CH2-- or --CH.dbd. and q is

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

SUMM . . . dexfenfluramine, dextroamphetamine, diethylpropion, diphemethoxidine, N-ethylamphetamine, fenbutrazate, fenfluramine, fenisorex, fenproporex, fludorex, fluminorex, furfurylmethylamphetamine, levamfetamine, levophacetoperane, mazindol, mefenorex, metamfepramone, methamphetamine, norpseudoephedrine, pentorex, phendimetrazine, phenmetrazine, phentermine, phenylpropanolamine, picilorex and sibutramine; and pharmaceutically acceptable salts thereof.

SUMM . . . as amphetamine, benzphetamine, chlorphentermine, clobenzorex, cloforex, clotermine, dexfenfluramine, dextroamphetamine, diethylpropion, N-ethylamphetamine, fenfluramine, fenproporex, furfurylmethylamphetamine, levamfetamine, mefenorex, metamfepramone, methamphetamine, norpseudoephedrine, pentorex, phendimetrazine, phenmetrazine, phentermine, phenylpropanolamine, picilorex and sibutramine; and pharmaceutically acceptable salts thereof.

SUMM . . . of obesity, such as, e.g., arteriosclerosis, Type II diabetes, polycycstic ovarian disease, cardiovascular diseases, osteoarthritis, dermatological disorders, hypertension, insulin resistance, hypercholesterolemia, hypertriglyceridemia, and cholelithiasis.

L42 ANSWER 21 OF 30 USPATFULL

ACCESSION NUMBER: 2001:52070 USPATFULL

TITLE: Substituted 3-(benzylamino)piperidine derivatives and

their use as therapeutic agents

INVENTOR(S): Elliott, Jason Matthew, Felsted, United Kingdom

PATENT ASSIGNEE(S): Merck Sharp & Dohme Limited, Hoddesdon, United States

(non-U.S. corporation)

WO 1998-GB1856 19980623

19991209 PCT 371 date 19991209 PCT 102(e) date

NUMBER DATE

PRIORITY INFORMATION: GB 1997-13715 19970627

GB 1997-20998 19971003

DOCUMENT TYPE: Utility
FILE SEGMENT: Granted
PRIMARY EXAMINER: Chang, Ceila

LEGAL REPRESENTATIVE: Thies, J. Eric, Rose, David L.

NUMBER OF CLAIMS: 9
EXEMPLARY CLAIM: 1
LINE COUNT: 1317

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

The present invention relates to compounds of formula (I), wherein R.sup.1 represents a fluoroC.sub.1-2 alkoxy group; and R.sup.2 represents a hydrogen or halogen atom or a C.sub.1-4 alkyl, C.sub.1-4 alkoxy, fluoroC.sub.1-4 alkyl or fluoroC.sub.1-4 alkoxy group; or a pharmaceutically acceptable salt thereof. The compounds are of particular use in the treatment or prevention of pain or inflammation, migraine, emesis, postherpetic neuralgia, depression or anxiety. ##STR1##

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

SUMM . . . dexfenfluramine, dextroamphetamine, diethylpropion, diphemethoxidine, N-ethylamphetamine, fenbutrazate, fenfluramine, fenisorex, fenproporex, fludorex, fluminorex, furfurylmethylamphetamine, levamfetamine, levophacetoperane, mazindol, mefenorex, metamfepramone, methamphetamine, norpseudoephedrine, pentorex, phendimetrazine, phenmetrazine, phentermine, phenylpropanolamine, picilorex and sibutramine; and pharmaceutically acceptable salts thereof.

SUMM . . . as amphetamine, benzphetamine, chlorphentermine, clobenzorex, cloforex, clotermine, dexfenfluramine, dextroamphetamine, diethylpropion, N-ethylamphetamine, fenfluramine, fenproporex, furfurylmethylamphetamine, levamfetamine, mefenorex, metamfepramone, methamphetamine, norpseudoephedrine, pentorex, phendimetrazine, phenmetrazine, phentermine, phenylpropanolamine, picilorex and sibutramine; and pharmaceutically acceptable salts thereof.

SUMM . . . of obesity, such as, e.g., arteriosclerosis, Type II diabetes, polycycstic ovarian disease, cardiovascular diseases, osteoarthritis, dermatological disorders, hypertension, insulin resistance,

hypercholesterolemia, hypertriglyceridemia, and cholelithiasis.

L42 ANSWER 22 OF 30 USPATFULL

ACCESSION NUMBER: 2001:44257 USPATFULL

TITLE: Pharmaceutical combinations for treating obesity and

food craving

INVENTOR(S): Rothman, Richard Brian, 8508 Carlynn Dr, Bethesda, MD,

United States 20817

DOCUMENT TYPE: Utility FILE SEGMENT: Granted

PRIMARY EXAMINER: Goldberg, Jerome D.

ASSISTANT EXAMINER: Kim, Jennifer

NUMBER OF CLAIMS: 4
EXEMPLARY CLAIM: 1
LINE COUNT: 433

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

Numerous studies have documented that medications which increase brain serotonin (5-HT) are effective anorectic agents which help obese patients lose weight and which also decrease craving for sweets and carbohydrates. Evidence from other studies also indicate that increases in brain 5-HT may help decrease craving for alcohol and cocaine.

5-hydroxy-L-tryptophan, abbreviated 5-HTP, is the immediate precursor of serotonin (5-HT). When administered in combination with an inhibitor of peripheral decarboxylase such as carbidopa, 5-HTP increases brain serotonin. Increases in synaptic 5-HT decreases the firing rate of 5-HT

neurons via stimulation of inhibitory 5-HT1a receptors located on the cell bodies in the raphe. This serves as a negative feedback loop. The clinically available beta adreneric receptor antagonist medication pindolol is also a 5-HT1a antagonist, and can be used to increase the ability of 5-HTP to increase brain 5-HT. Previous studies with 5-HTP used doses exceeding 50 mg per day. When 5-HTP was used in combination with carbidopa, the dose of carbidopa was in excess of 50 mg per day. One novel aspect of the invention are the doses of the 5-HTP and carbidopa: much lower daily doses than have been used before are effective in decreasing appetite, decreasing craving for food and for promoting weight loss. The second novel aspect of the invention relates to the concurrent use of pindolol along with the 5-HTP/Carbidopa, which enhances the effectiveness of the 5-HTP/Carbidopa combination.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

DETD . . . the treatment of diseases which are complications to overweight or obesity. These diseases or conditions include diabetes mellitus type II, hypercholesterolemia, hypertriglyceridaemia, hypertension, back pain caused by obesity, arthritis made worse by obesity, sleep apnea and psychological or psychiatric problems complicated. . .

DETD In a further aspect, the invention can be used along with other appetite suppressant drugs, such as amphetamine, phentermine, diethylpropion, phendimetrazine, ephedrine or similarly-acting agents, to enhance the effects of these medications, that is to increase the weight loss which would. . .

CLM What is claimed is:

. also be administered in combination with an effective amount of other anorectic agents selected from group consisting of phentermine, diethylpropion, phendimetrazine and ephedrine.

IT 50-67-9, Serotonin, biological studies 90-84-6, Diethylpropion
122-09-8, Phentermine 299-42-3, Ephedrine 634-03-7,
Phendimetrazine 4350-09-8, L-5-Hydroxytryptophan 13523-86-9, Pindolol
28860-95-9, Carbidopa

(antiobesity compns. contg. synergistic combination of L-5-hydroxytryptophan and carbidopa and pindolol and other agents)

=> log h SINCE FILE COST IN U.S. DOLLARS TOTAL ENTRY SESSION FULL ESTIMATED COST 18.28 225.98 DISCOUNT AMOUNTS (FOR QUALIFYING ACCOUNTS) SINCE FILE TOTAL ENTRY SESSION CA SUBSCRIBER PRICE 0.00 -6.51

SESSION WILL BE HELD FOR 60 MINUTES
STN INTERNATIONAL SESSION SUSPENDED AT 11:33:35 ON 08 JAN 2003

Welcome to STN International! Enter x:x

LOGINID:sssptal617srh

PASSWORD:

TERMINAL (ENTER 1, 2, 3, OR ?):2

NEWS 1 Web Page URLs for STN Seminar Schedule - N. America
NEWS 2 Apr 08 "Ask CAS" for self-help around the clock
NEWS 3 Apr 09 BEILSTEIN: Reload and Implementation of a New Subject Area
NEWS 4 Apr 09 ZDB will be removed from STN
NEWS 5 Apr 19 US Patent Applications available in IFICDB, IFIPAT, and IFIUDB
NEWS 6 Apr 22 Records from IP.com available in CAPLUS, HCAPLUS, and ZCAPLUS
NEWS 7 Apr 22 BIOSIS Gene Names now available in TOXCENTER
NEWS 8 Apr 22 Federal Research in Progress (FEDRIP) now available
NEWS 9 Jun 03 New e-mail delivery for search results now available
NEWS 10 Jun 10 MEDLINE Reload
NEWS 11 Jun 10 PCTFULL has been reloaded

```
NEWS 12 Jul 02 FOREGE no longer contains STANDARDS file segment
 NEWS 13 Jul 22 USAN to be reloaded July 28, 2002;
                 saved answer sets no longer valid
 NEWS 14 Jul 29
                 Enhanced polymer searching in REGISTRY
 NEWS 15 Jul 30 NETFIRST to be removed from STN
 NEWS 16 Aug 08
                 CANCERLIT reload
 NEWS 17 Aug 08
                 PHARMAMarketLetter(PHARMAML) - new on STN
 NEWS 18 Aug 08 NTIS has been reloaded and enhanced
                 Aquatic Toxicity Information Retrieval (AQUIRE)
 NEWS 19 Aug 19
                 now available on STN
 NEWS 20 Aug 19
                 IFIPAT, IFICDB, and IFIUDB have been reloaded
 NEWS 21 Aug 19
                 The MEDLINE file segment of TOXCENTER has been reloaded
                 Sequence searching in REGISTRY enhanced
 NEWS 22 Aug 26
 NEWS 23 Sep 03
                 JAPIO has been reloaded and enhanced
 NEWS 24 Sep 16
                 Experimental properties added to the REGISTRY file
 NEWS 25 Sep 16
                 Indexing added to some pre-1967 records in CA/CAPLUS
 NEWS 26 Sep 16 CA Section Thesaurus available in CAPLUS and CA
 NEWS 27 Oct 01
                 CASREACT Enriched with Reactions from 1907 to 1985
 NEWS 28 Oct 21 EVENTLINE has been reloaded
 NEWS 29 Oct 24 BEILSTEIN adds new search fields
 NEWS 30 Oct 24 Nutraceuticals International (NUTRACEUT) now available on STN
 NEWS 31 Oct 25 MEDLINE SDI run of October 8, 2002
NEWS 32 Nov 18 DKILIT has been renamed APOLLIT
 NEWS 33 Nov 25 More calculated properties added to REGISTRY
 NEWS 34 Dec 02 TIBKAT will be removed from STN
 NEWS 35 Dec 04 CSA files on STN
 NEWS 36 Dec 17 PCTFULL now covers WP/PCT Applications from 1978 to date
 NEWS 37 Dec 17 TOXCENTER enhanced with additional content
 NEWS 38 Dec 17 Adis Clinical Trials Insight now available on STN
NEWS 39 Dec 30 ISMEC no longer available
NEWS EXPRESS January 6 CURRENT WINDOWS VERSION IS V6.01a,
              CURRENT MACINTOSH VERSION IS V6.0b(ENG) AND V6.0Jb(JP),
              AND CURRENT DISCOVER FILE IS DATED 01 OCTOBER 2002
NEWS HOURS
              STN Operating Hours Plus Help Desk Availability
 NEWS INTER
              General Internet Information
NEWS LOGIN
              Welcome Banner and News Items
NEWS PHONE
              Direct Dial and Telecommunication Network Access to STN
NEWS WWW
              CAS World Wide Web Site (general information)
Enter NEWS followed by the item number or name to see news on that
specific topic.
 All use of STN is subject to the provisions of the STN Customer
 agreement. Please note that this agreement limits use to scientific
 research. Use for software development or design or implementation
 of commercial gateways or other similar uses is prohibited and may
 result in loss of user privileges and other penalties.
  FILE 'HOME' ENTERED AT 12:36:04 ON 08 JAN 2003
=> d his
     (FILE 'HOME' ENTERED AT 12:36:04 ON 08 JAN 2003)
=> fil reg
COST IN U.S. DOLLARS
                                               SINCE FILE
                                                              TOTAL
                                                    ENTRY
                                                            SESSION
FULL ESTIMATED COST
                                                     0.21
                                                               0.21
```

FILE 'REGISTRY' ENTERED AT 12:36:22 ON 08 JAN 2003 USE IS SUBJECT TO THE TERMS OF YOUR STN CUSTOMER AGREEMENT. PLEASE SEE "HELP USAGETERMS" FOR DETAILS. COPYRIGHT (C) 2003 American Chemical Society (ACS)

Property values tagged with IC are from the ZIC/VINITI data file provided by InfoChem.

STRUCTURE FILE UPDATES: 7 JAN 2003 HIGHEST RN 478336-86-6 DICTIONARY FILE UPDATES: 7 JAN 2003 HIGHEST RN 478336-86-6

```
TSCA INFORMATION NOW CURRENT THROUGH MAY 20, 2002
```

Please note that search-term pricing does apply when conducting SmartSELECT searches.

Crossover limits have been increased. See HELP CROSSOVER for details.

Experimental and calculated property data are now available. See HELP PROPERTIES for more information. See STNote 27, Searching Properties in the CAS Registry File, for complete details: http://www.cas.org/ONLINE/STN/STNOTES/stnotes27.pdf

=> d

L1 ANSWER 1 OF 1 REGISTRY COPYRIGHT 2003 ACS

RN 634-03-7 REGISTRY

CN Morpholine, 3,4-dimethyl-2-phenyl-, (2S,3S)- (9CI) (CA INDEX NAME)

OTHER CA INDEX NAMES:

CN Morpholine, 3,4-dimethyl-2-phenyl-, (+)- (8CI)

CN Morpholine, 3,4-dimethyl-2-phenyl-, (2S-trans)-

THER NAMES:

CN (+)-3,4-Dimethyl-2-phenylmorpholine

CN (+)-Phendimetrazine

CN Antapentan

CN d-2-Phenyl-3,4-dimethylmorpholine

CN Mephenmetrazine

CN Phendimetrazine

CN Sedafamen

FS STEREOSEARCH

DR 711-71-7, 16311-86-7

MF C12 H17 N O

CI COM

LC STN Files: ADISNEWS, ANABSTR, BEILSTEIN*, BIOSIS, BIOTECHNO, CA, CAOLD, CAPLUS, CBNB, CHEMLIST, CIN, DDFU, DIOGENES, DRUGU, EMBASE, HSDB*, IPA, MRCK*, RTECS*, SPECINFO, TOXCENTER, USAN, USPAT2, USPATFULL (*File contains numerically searchable property data)

Other Sources: EINECS**, WHO

(**Enter CHEMLIST File for up-to-date regulatory information)

Absolute stereochemistry.

PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT

177 REFERENCES IN FILE CA (1962 TO DATE)
177 REFERENCES IN FILE CAPLUS (1962 TO DATE)
3 REFERENCES IN FILE CAOLD (PRIOR TO 1967)

=> sel rn name

E1 THROUGH E8 ASSIGNED

=> fil medl capl biosis uspatf COST IN U.S. DOLLARS

SINCE FILE TOTAL ENTRY SESSION

FULL ESTIMATED COST

6.67 6.88

FILE 'CAPLUS' ENTERED AT 12:36:54 ON 08 JAN 2003 USE IS SUBJECT TO THE TERMS OF YOUR STN CUSTOMER AGREEMENT. PLEASE SEE "HELP USAGETERMS" FOR DETAILS. COPYRIGHT (C) 2003 AMERICAN CHEMICAL SOCIETY (ACS) FILE 'BIOSIS' ENTERED AT 12:36:54 ON 08 JAN 2003 COPYRIGHT (C) 2003 BIOLOGICAL ABSTRACTS INC. (R) FILE 'USPATFULL' ENTERED AT 12:36:54 ON 08 JAN 2003 CA INDEXING COPYRIGHT (C) 2003 AMERICAN CHEMICAL SOCIETY (ACS)

L2

463 ("(+)-PHENDIMETRAZINE"/BI OR "(+)-3,4-DIMETHYL-2-PHENYLMORPHOLIN E"/BI OR ANTAPENTAN/BI OR "D-2-PHENYL-3,4-DIMETHYLMORPHOLINE"/BI OR MEPHENMETRAZINE/BI OR PHENDIMETRAZINE/BI OR SEDAFAMEN/BI OR 634-03-7/BI)

=> s weight control or weight loss or weight gain or obesity or obese or fat 795810 WEIGHT CONTROL OR WEIGHT LOSS OR WEIGHT GAIN OR OBESITY OR OBESE OR FAT

=> s 12 and 13

L4118 L2 AND L3

=> s hypercholesterol? or hypocholesterolem?

59667 HYPERCHOLESTEROL? OR HYPOCHOLESTEROLEM?

=> s 14 and 15

33 L4 AND L5

=> dup rem 16

PROCESSING COMPLETED FOR L6

33 DUP REM L6 (0 DUPLICATES REMOVED)

=> d ibib abs 20-23

1.7 ANSWER 20 OF 33 USPATFULL

2001:212586 USPATFULL ACCESSION NUMBER:

TITLE: In vivo delivery methods and compositions INVENTOR(S): Kensey, Kenneth R., Malvern, PA, United States

NUMBER KIND DATE -----PATENT INFORMATION: US 2001044584 A1 20011122 US 2001-819924 APPLICATION INFO.: A1 20010328 (9)

RELATED APPLN. INFO.: Continuation-in-part of Ser. No. US 2000-727950, filed on 1 Dec 2000, PENDING Continuation-in-part of Ser. No.

US 2000-628401, filed on 1 Aug 2000, PENDING

Continuation-in-part of Ser. No. US 2000-501856, filed on 10 Feb 2000, PENDING Continuation-in-part of Ser. No. US 1999-439795, filed on 12 Nov 1999, PENDING Continuation-in-part of Ser. No. US 1997-919906, filed

on 28 Aug 1997, GRANTED, Pat. No. US 6019735

DOCUMENT TYPE: Utility

FILE SEGMENT: APPLICATION

LEGAL REPRESENTATIVE: CAESAR, RIVISE, BERNSTEIN,, COHEN & POKOTILOW, LTD., 12TH FLOOR, SEVEN PENN CENTER, 1635 MARKET STREET,

PHILADELPHIA, PA, 19103-2212

NUMBER OF CLAIMS: 36

1

EXEMPLARY CLAIM: NUMBER OF DRAWINGS:

19 Drawing Page(s)

2120

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

Various methods are provided for determining and utilizing the viscosity of the circulating blood of a living being over a range of shear rates for diagnostics and treatment, such as detecting/reducing blood viscosity, work of the heart, contractility of the heart, for detecting/reducing the surface tension of the blood, for detecting plasma viscosity, for explaining/countering endothelial cell dysfunction, for providing high and low blood vessel wall shear stress data, red blood cell deformability data, lubricity of blood, and for

treating different ailments such as peripheral arterial disease in combination with administering to a living being at least one pharmaceutically acceptable agent. Agents pharmaceutically effective to regulate at least one of the aforementioned blood parameters are used to adjust distribution of a substance through the bloodstream.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

L7 ANSWER 21 OF 33 USPATFULL

ACCESSION NUMBER: 2001:188226 USPATFULL

TITLE: DIETETIC FOOD COMPOSITION AND DIETETIC METHOD USING

SUCH COMPOSITION

INVENTOR(S): ZOHOUNGBOGBO, MATHIAS CHRISTIAN, RIVALTA DI TORINO,

Italy

RELATED APPLN. INFO.: Continuation-in-part of Ser. No. US 1999-225819, filed

on 5 Jan 1999, ABANDONED

DOCUMENT TYPE: Utility
FILE SEGMENT: APPLICATION

LEGAL REPRESENTATIVE: SOFER & HAROUN LLP, 342 MADISON AVENUE, SUITE 1921, NEW

YORK, NY, 10173

NUMBER OF CLAIMS: 42 EXEMPLARY CLAIM: 1 LINE COUNT: 833

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

AB Food composition in the form of a flour comprising at least 50% of protein, less than 15% of carbohydrates and 35 to 50% of plant fibers; preferably the carbohydrate content is less than 10%, advantageously less than 5%; this composition may be used as a substitute for wheat flour in the preparation of foods such as pasta, bread, bread sticks, bakery products and pastries and constitutes the basis of a method for improving the appearance of a person by achieving a loss of weight which is beneficial from the aesthetic point of view.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

L7 ANSWER 22 OF 33 USPATFULL

ACCESSION NUMBER: 2001:90260 USPATFULL

TITLE: Fatty acid-pharmaceutical agent conjugates INVENTOR(S): Webb, Nigel L., Bryn Mawr, PA, United States

Bradley, Matthews O., Laytonsville, MD, United States Swindell, Charles S., Merion, PA, United States Shashoua, Victor E., Brookline, MA, United States

APPLICATION INFO.: US 2000-730450 A1 20001205 (9)
RELATED APPLN. INFO.: Continuation of Ser. No. US 1996-651428, filed on 22

May 1996, ABANDONED

DOCUMENT TYPE: Utility FILE SEGMENT: APPLICATION

LEGAL REPRESENTATIVE: Edward R. Gates, Wolf, Greenfield & Sacks, P.C., 600

Atlantic Avenue, Boston, MA, 02210

NUMBER OF CLAIMS: 12 EXEMPLARY CLAIM: 1

PATENT INFORMATION:

NUMBER OF DRAWINGS: 14 Drawing Page(s)

LINE COUNT: 2511

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

AB The invention provides conjugates of fatty acids and pharmaceutical agents useful in treating noncentral nervous system conditions. Methods for selectively targeting pharmaceutical agents to desired tissues are provided.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

L7 ANSWER 23 OF 33 USPATFULL

ACCESSION NUMBER: 2001:63697 USPATFULL

TITLE: Spiro-azacyclic derivatives and their use as

therapeutic agents

INVENTOR(S): Kulagowski, Janusz Jozef, Sawbridgeworth, United

Kingdom

Raubo, Piotr Antoni, Bishops Stortford, United Kingdom Swain, Christopher John, Duxford, United Kingdom Thomson, Christopher George, Sawbridgeworth, United

Kingdom

PATENT ASSIGNEE(S): Merck Sharp & Dohme Ltd., Hertfordshire, United Kingdom

(non-U.S. corporation)

19991118 PCT 371 date 19991118 PCT 102(e) date

NUMBER DATE

PRIORITY INFORMATION: GB 1997-11114 19970529

DOCUMENT TYPE: Utility FILE SEGMENT: Granted

PRIMARY EXAMINER: Rotman, Alan L.
ASSISTANT EXAMINER: Desai, Rita

LEGAL REPRESENTATIVE: Thies, J. Eric, Rose, David L.

NUMBER OF CLAIMS: 19
EXEMPLARY CLAIM: 1
LINE COUNT: 3494

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

AB Substituted spiro-azacyclic derivatives of structural formula I are tachykinin receptor antagonists of use, for example, in the treatment of pain, inflammation, migraine, emesis and posttherpetic neuralgia ##STR1##

Wherein A is a pyridyl, X is --CH2--, Y is --CH2-- or --CH.dbd. and q is 2.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

=> d ibib abs 30-33

L7 ANSWER 30 OF 33 USPATFULL

ACCESSION NUMBER: 1998:98932 USPATFULL

TITLE: DHA-pharmaceutical agent conjugates of taxanes
INVENTOR(S): Shashoua, Victor E., Brookline, MA, United States
Swindell, Charles S., Merion, PA, United States
Webb, Nigel L., Bryn Mawr, PA, United States

Bradley, Matthews O., Laytonsville, MD, United States PATENT ASSIGNEE(S): Neuromedica, Inc., Conshohocken, PA, United States

(U.S. corporation)

DOCUMENT TYPE: Utility
FILE SEGMENT: Granted

PRIMARY EXAMINER: Jarvis, William R. A.

LEGAL REPRESENTATIVE: Wolf, Greenfield & Sacks, P.C.

NUMBER OF CLAIMS: 12 EXEMPLARY CLAIM: 1

NUMBER OF DRAWINGS: 27 Drawing Figure(s); 14 Drawing Page(s)

LINE COUNT: 2451

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

The invention provides conjugates of cis-docosahexaenoic acid and taxanes useful in treating cell proliferative disorders. Conjugates of paclitaxel and docetaxel are preferred.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

ANSWER 31 OF 33 USPATFULL

ACCESSION NUMBER: 1998:98904 USPATFULL

TITLE: Method and composition for treating obesity

and related disorders in animals comprising

dehydroepiandrosterone (DHEA), or a derivative thereof,

and an anorectic agent

Svec, Frank, Metairie, LA, United States INVENTOR(S): Porter, Johnny, Metairie, LA, United States

PATENT ASSIGNEE(S): Louisiana State University Medical Center Foundation,

New Orleans, LA, United States (U.S. corporation)

NUMBER KIND DATE _____ PATENT INFORMATION: US 5795880 19980818 APPLICATION INFO.: US 1996-774521 19961230 (8) DOCUMENT TYPE: Utility

FILE SEGMENT: Granted

PRIMARY EXAMINER: Weddington, Kevin E.

LEGAL REPRESENTATIVE: Finnegan, Henderson, Farabow, Garrett & Dunner, L.L.P.

NUMBER OF CLAIMS: 37 EXEMPLARY CLAIM:

NUMBER OF DRAWINGS: 27 Drawing Figure(s); 13 Drawing Page(s) LINE COUNT: 888

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

The invention describes a method and composition for treating obesity or related disorders in animals using an anorectic agent and dehydroepiandrosterone (DHEA). The composition effectively

diminishes caloric intake, may alter metabolism, weight

gain, or a combination thereof.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

ANSWER 32 OF 33 USPATFULL

ACCESSION NUMBER: 97:7901 USPATFULL

Method for treatment or prevention of obesity TITLE: INVENTOR (S): Clark, Ross G., Pacifica, CA, United States

PATENT ASSIGNEE(S): Genentech, Inc., South San Francisco, CA, United States

(U.S. corporation)

NUMBER KIND DATE US 5597797 19970128 PATENT INFORMATION: WO 9118621 19911212 19931119 (8) 19931026 APPLICATION INFO.: US 1993-150090 WO 1993-US10259

19931119 PCT 371 date 19931119 PCT 102(e) date

DOCUMENT TYPE: Utility FILE SEGMENT: Granted

PRIMARY EXAMINER: Schain, Howard E. ASSISTANT EXAMINER: Touzeau, P. Lynn LEGAL REPRESENTATIVE: Hasak, Janet E.

NUMBER OF CLAIMS: 20 EXEMPLARY CLAIM: 1

NUMBER OF DRAWINGS: 20 Drawing Figure(s); 20 Drawing Page(s)

LINE COUNT: 2197

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

A method is disclosed for treating obese mammals or preventing obesity from occurring in mammals. This method involves administering to the mammal an effective amount of growth hormone in combination with an effective amount of IGF-I. Preferably, the growth hormone is given so as to have a maintained, continual therapeutically effective presence in the blood, such as by continuous infusion or frequent injections, or by use of a long-acting formulation.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

L7 ANSWER 33 OF 33 USPATFULL

ACCESSION NUMBER:

96:53303 USPATFULL

TITLE:

Method and composition for treating obesity comprising dehydroepiandrosterone (DHEA), or a derivative thereof, and an anorectic agent

INVENTOR (S):

Svec, Frank, Metairie, LA, United States Porter, Johnny, Metairie, LA, United States

PATENT ASSIGNEE(S):

Louisiana State Univ. Medical Center Foundation, New

Orleans, LA, United States (U.S. corporation)

NUMBER KIND DATE -----

PATENT INFORMATION:

US 5527788

19960618

APPLICATION INFO.:

US 1994-184191

19940118 (8)

DOCUMENT TYPE: FILE SEGMENT:

Utility Granted

PRIMARY EXAMINER:

Cintins, Marianne M.

ASSISTANT EXAMINER:

Weddington, Kevin E.

LEGAL REPRESENTATIVE: Finnegan, Henderson, Farabow, Garrett & Dunner

NUMBER OF CLAIMS: EXEMPLARY CLAIM:

NUMBER OF DRAWINGS:

1 27 Drawing Figure(s); 10 Drawing Page(s)

LINE COUNT:

799

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

The invention describes a method and composition for treating obesity or related disorders in animals using an anorectic agent and dehydroepiandrosterone (DHEA). The composition effectively diminishes caloric intake, may alter metabolism, weight

gain, or a combination thereof.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

=> fil stng

COST IN U.S. DOLLARS

SINCE FILE TOTAL

ENTRY 113.66 SESSION 120.54

FULL ESTIMATED COST

FILE 'STNGUIDE' ENTERED AT 13:06:48 ON 08 JAN 2003 USE IS SUBJECT TO THE TERMS OF YOUR CUSTOMER AGREEMENT COPYRIGHT (C) 2003 AMERICAN CHEMICAL SOCIETY, JAPAN SCIENCE AND TECHNOLOGY CORPORATION, AND FACHINFORMATIONSZENTRUM KARLSRUHE

FILE CONTAINS CURRENT INFORMATION.

LAST RELOADED: Dec 20, 2002 (20021220/UP).

---Logging off of STN---

Executing the logoff script...

=> LOG Y

COST IN U.S. DOLLARS

SINCE FILE

FULL ESTIMATED COST

SESSION ENTRY 1.62 122.16

TOTAL

STN INTERNATIONAL LOGOFF AT 13:23:00 ON 08 JAN 2003

```
AN
     93239105
                  MEDLINE
                PubMed ID: 8477962
DN
     93239105
     Cholesterol-lowering effect of ursodeoxycholic acid in patients with
TI
     primary biliary cirrhosis.
ΑU
     Poupon R E; Ouquerram K; Chretien Y; Verneau C; Eschwege E; Magot T;
     Poupon R
     INSERM U21, Unite de Recherches Cliniques et Epidemiologiques, 94807
CS
     Villejuif, France.
SO
     HEPATOLOGY, (1993 Apr) 17 (4) 577-82.
     Journal code: 8302946. ISSN: 0270-9139.
     United States
CY
DT
     (CLINICAL TRIAL)
     Journal; Article; (JOURNAL ARTICLE)
     (RANDOMIZED CONTROLLED TRIAL)
LA
     English
FS
     Priority Journals
EΜ
     199305
ED
     Entered STN: 19930611
     Last Updated on STN: 19930611
     Entered Medline: 19930525
AB
     We have previously shown in a 2-yr controlled trial that
     hypercholesterolemia, frequent in primary biliary cirrhosis, is
     lowered by ursodeoxycholic acid (13 to 15 mg daily).
     To further investigate this effect, we analyzed the influence of long-term
     ursodeoxycholic acid administration on serum lipids,
     lipoproteins and bile acids. The study involved a subgroup of 33
     noncirrhotic patients (17 received ursodeoxycholic acid
     and 16 received a placebo) analyzed at inclusion and after 2 yr. The total
     serum cholesterol concentration was markedly reduced in the
    ursodeoxycholic acid-treated patients in comparison with
     the controls (mean +/- S.E.M. = 7.49 +/- 0.42 mmol/L and 7.07 +/- 0.23
    mmol/L at entry and 4.44 +/- 0.40 mmol/L and 6.89 +/- 0.27 mmol/L at 2 yr
     in the ursodeoxycholic acid and placebo groups,
     respectively; p < 0.02). Quantitatively, this decrease was mainly caused
    by a fall in low-density-lipoprotein cholesterol, but very low
    density-lipoprotein cholesterol levels also fell significantly.
    High-density-lipoprotein cholesterol levels remained stable in both
    groups, but the high-density-lipoprotein2/high-density-lipoprotein3
    cholesterol ratio fell significantly during ursodeoxycholic
     acid treatment. No significant change occurred in total
     triglyceride or total phospholipid levels. In the treated group, the
    proportion of ursodeoxycholic acid increased (up to
    60% of total circulating bile acids), whereas that of cholic and
    chenodeoxycholic acids fell significantly. In conclusion, the
    cholesterol-lowering effect of ursodeoxycholic acid
    could be related to an improvement of cholestasis, modifications in
    cholesterol metabolism or both. Changes in endogenous bile acid
    composition induced by ursodeoxycholic acid might be
    the common denominator of these two mechanisms.
    Check Tags: Comparative Study; Human
     *Anticholesteremic Agents: TU, therapeutic use
     *Bile Acids and Salts: BL, blood
     *Cholesterol: BL, blood
     Double-Blind Method
     Follow-Up Studies
     Lipoproteins, HDL Cholesterol: BL, blood
     Lipoproteins, LDL Cholesterol: BL, blood
     *Liver Cirrhosis, Biliary: BL, blood
     *Liver Cirrhosis, Biliary: DT, drug therapy
     Liver Function Tests
     Middle Age
```

Placebos
Time Factors
*Ursodeoxycholic Acid: TU, therapeutic use
RN 128-13-2 (Ursodeoxycholic Acid); 57-88-5 (Cholesterol)
CN 0 (Anticholesteremic Agents); 0 (Bile Acids and Salts); 0 (Lipoproteins, HDL Cholesterol); 0 (Lipoproteins, LDL Cholesterol); 0 (Placebos)

ACCESSION NUMBER: 2000220279 MEDLINE

DOCUMENT NUMBER: 20220279 PubMed ID: 10756780

TITLE: [The importance of the use of selenium in the role of an

antioxidant in preventing cardiovascular diseases]. Importanta utilizarii seleniului cu rol antioxidant in

preventia bolilor cardiovasculare.

AUTHOR: Azoicai D; Ivan A; Bradatean M; Pavel M; Jerca L;

Iacobovici A; Popovici I; Gheorghita N

CORPORATE SOURCE: Disciplina de Epidemiologie, Facultatea de Medicina,

Universitatea de Medicina si Farmacie Gr. T. Popa, Iasi. REVISTA MEDICO-CHIRURGICALA A SOCIETATII DE MEDICI SI

NATURALISTI DIN IASI, (1997 Jul-Dec) 101 (3-4) 109-15.

Journal code: 0413735. ISSN: 0300-8738.

PUB. COUNTRY: Romania

SOURCE:

DOCUMENT TYPE: Journal; Article; (JOURNAL ARTICLE)

LANGUAGE: Romanian

FILE SEGMENT: Priority Journals

ENTRY MONTH: 200005

ENTRY DATE: Entered STN: 20000606

Last Updated on STN: 20000606 Entered Medline: 20000519

The evaluation of the results of the oxygen free radicals (RLO2) formation AB is a current subject in biology and medicine. The oxidative stress, which is the altering of the balance between the higher activity of oxygen and the enzymatic or nonenzymatic protection systems, may be one of the causes that starts and aggravates a disease. In this context, the supplementation of the diet with selenium, superoxide dismutase, vitamins A, C, E, is considered a primary prevention measure (for the apparently healthy persons) and a secondary one (for those with advancing forms of disease) that is both efficient and modern by utilization of some "drug-food" products. The transversal study realized on a group of 39 blood donors presence of the cardiovascular risk determined by the raising of the prevalence of some atherogenic factors (active smoking, hypercholesterolemia) which is also expressed by the lowering of the level of some oxidative stress indicators (glutathione peroxidase--GSH-Px < 0.139 moli/ml and catalase < 2.20 U/ml). The simultaneous low intake of selenium from the central drinking water supplies in the city of Iasi (0.1-1 q/l) has permitted us to consider necessary the diet supplementation both with foods rich in vitamins with an antioxidant role and with specific medication with selenium, as a protective micro-element.

ACCESSION NUMBER: 80025910 MEDLINE

DOCUMENT NUMBER: 80025910 PubMed ID: 488876

TITLE: [Possibilities for weight reduction by means of diet].

Moglichkeiten zur Verminderung des Korpergewichts mittels

diatetischer Massnahmen.

AUTHOR: Forster H

SOURCE: FORTSCHRITTE DER MEDIZIN, (1979 Aug 23) 97 (32) 1339-44.

Journal code: 2984763R. ISSN: 0015-8178.

PUB. COUNTRY: GERMANY, WEST: Germany, Federal Republic of

DOCUMENT TYPE: Journal; Article; (JOURNAL ARTICLE)

LANGUAGE: German

FILE SEGMENT: Priority Journals

ENTRY MONTH: 197912

ENTRY DATE: Entered STN: 19900315

Last Updated on STN: 19900315 Entered Medline: 19791218

AB The different dietetic measures for weight reduction are described. According to the existing overweight the therapeutic measures are classified in four steps. In the first step, with low overweight, the energy-containing drinks (soft drinks and alcoholic beverages) are avoided. If the overweight is greater an additional reduction of the energy content of meal is required. A real reduction-diet (less than 1.000 Kcal/day or 4.200 KJ/day) demands extensive knowledge of food composition and greater efforts in meal composition. The availability of formula diets is considered as a relief. During starvation (or total fasting) as the step 4 of weight reduction diet, an extreme metabolic alteration takes place, which is characterized by ketosis. The same metabolic alteration is found by a fat-protein-diet (a so-called ketogenic diet), where hypercholesterolemia and hyperuricemia are common side effects. The carbohydrate-protein weight reduction diet is poor in health risks. Furthermore the normal metabolic pattern is maintained during this kind of diet if enough carbohydrates are provided per day (i.e. 80-100 g/day).